

Anti-HB Ag Immunoglobulin Staves Off Clinical Hepatitis B

BY JAMES MAGEE
Medical Tribune Staff

MILAN, ITALY—A British team has reported successful prevention of the appearance of clinical hepatitis B following accidental inoculations with antigen bearing material, through prophylactic use of anti-HB Ag immunoglobulin.

Reporting the first results of an uncontrolled Medical Research Council trial involving 110 persons with inoculation injuries, Dr. Sheila Polakov said the average interval from accident to prophylaxis was six days. More than 80 per cent of the participants were given prophylaxis within eight days of the accident. The trial began in September 1973.

2 of 61 Develop Jaundice

Accidents included transfusion of blood or blood products, penetration of the skin, contamination of the conjunctival sac or cuts or abrasions of the skin involving material containing HB Ag, whether blood, other body fluids, or laboratory reagents, she told a symposium on viral hepatitis at the International Association of Biological Standardization meeting here.

Of 61 study participants whose accidents involved penetration, and who were followed, to date two have developed jaundice with hepatitis B antigenemia. In one the illness began 17 weeks after the accident. The attack was mild and the antigenemia transient: the patient made a full clinical recovery within five weeks of the onset.

The first indication of illness in the other was detection of HB Ag and raised aminotransferase levels in a follow-up sample taken 18 weeks after the accident: jaundice and other manifestations developed shortly afterwards. The attack was recent and the course of the illness is still being monitored.

4 Globulin Studies On in US

[Currently, four double-blind studies are underway in the United States to test the efficacy of hyperimmune globulin in preventing clinical hepatitis B.

[Three of these studies are under the direction of the National Heart and Lung Institute's Division of Blood Diseases and Resources, including trials of the globulin in renal dialysis patients, patients with needle stick and other accidental exposure, and transfused patients. The VA also has a double-blind study underway.

[Dr. Harvey G. Klein, project officer for the NHLI studies, noted that if early results indicate the hyperimmune globulin is effective, a Data Safety Monitoring Committee headed by Douglas M. Surgeon, Ph.D., former dean of the State University of New York's Buffalo School of Medicine, will halt the study and release the data.]

Dr. Polakov, who is associated with the epidemiological research laboratory, Central Public Health Laboratory, London, explained that when an accident that meets the study criteria is reported a sample of the inoculated material is tested for HB Ag. If a sample is not available, documentary

evidence of the presence of HB Ag by previous tests of the material or, if the source is a person, of samples taken at any time in the four weeks before the accident, is accepted.

A serum sample, taken from the person who sustained the accident, is also tested by routine methods for HB Ag and anti-HB Ag. If the results of these tests are negative and the immunoglobulin can be administered within approximately two weeks of the accident, the subject is enrolled in the study. A 500 mg. dose of the material is given intramuscularly and the subject is observed for any immediate reactions.

Each subject is followed-up for one year after the accident. Serum samples are taken, usually at four weekly intervals in the first six months; two further samples are taken, one at nine months and the other at or about one year after the accident.

In the first year of study 110 persons were enrolled. Most of the participants were nursing, medical or laboratory staff: two were patients who had been infused with a blood product later found to be contaminated. Of the 110 accidents, penetration of

the skin accounted for more than half. There was no evidence of infection among subjects who contaminated abrasions or ingested infected material.

Anti-HB Ag Detected in 3

None of the participants appears to have developed asymptomatic HB Ag carriage, but anti-HB Ag was detected by counterimmunoelectrophoresis in sera from three, who had no other evidence of infection, at 18, 20 and 23 weeks after the accident. In one case anti-HB Ag was transitory; in another it is still present 20 weeks after it was first detected; in the third case it was detected in the most recent specimen. Four subjects each had a notably raised aminotransferase level in one follow-up specimen—one at 14 weeks, two at approximately 20 weeks and one at 27 weeks after the accident: none of the four had any other evidence of infection.

"These are of course preliminary results: further laboratory tests which will be made at the end of the study may reveal evidence of infection that could not be detected by the test methods in routine use," Dr. Polakov concluded.

Co-author was Dr. W. d'A. Maycock, The Lister Institute of Preventive Medicine, Elstree, Herts, United Kingdom.

Laparoscopy 'Best' of 4 Sterilization Routes

Medical Tribune World Service

Buenos Aires—Laparoscopy appears to be superior to colpotomy, culdoscopy, or laparotomy for sterilization of women who have not recently been pregnant, according to a study by the International Fertility Research Program (IFRP).

Complications of the four procedures during surgery and in the first to eighth postoperative weeks were reported at the Eighth World Congress on Fertility and Sterility by Dr. William E. Brenner and David A. Edelman, Ph.D.

They evaluated the results of 401 culdoscopies, 799 colpotomies, 482 laparoscopies, and 279 laparotomies performed at 11 American institutions from October, 1972, to December, 1973.

The most common operative difficulty with endoscopic methods was inadequate visualization of the tubes. This occurred in 3.5 per cent of culdoscopies and 2.5 per cent of laparoscopies.

Blood loss greater than 100 ml was more common with both vaginal methods.

Postoperative pelvic infections were more frequent with the vaginal methods—6 per cent with culdoscopy and 4.5 per cent with colpotomy.

Incision complications were more common with the abdominal approaches.

Operative and hospitalization times were significantly shorter with the endoscopic methods, and the proportion of women resuming normal activities within four weeks of sterilization was higher.

While technical difficulties, operative complications, surgical and hospitalization times, and resumption of activities were similar with laparos-

copy and culdoscopy, pelvic infection was more common with culdoscopy.

Dr. Brenner is director of IFRP research and training and Associate Professor at the University of North Carolina. Dr. Edelman is on the staff of the University of North Carolina.

Treatment of Sterility

► A Japanese physician reported that of 100 sterile women treated with clomiphene citrate, ovulation was induced in 84 and 39 became pregnant.

Dr. Tarao Shimomura, of Kitano Hospital, Osaka, said that the patients included 65 with primary sterility.

Thirteen of the 100 patients complained of infrequent ovulation; 26, anovulatory menstruation; 55, first-grade amenorrhea, which responded to

progesterone, with bleeding; and six,

Thin Fiberscope Facilitates Studies Of Esophagus, Stomach, Duodenum

Medical Tribune World Service

Mexico City—In endoscopic examination of the esophagus, stomach, and duodenum, a fiberscope that is about half the standard size has shown significant advantages, it was reported at the Third International Congress of Gastrointestinal Endoscopy.

This instrument, Olympus GIF-P, with a tube diameter of 6.8 mm can be passed with little premedication. It was originally developed for esophageal cancer surveying in Japan, and has been widely employed there. Initially, it was brought to the United States for pediatric endoscopy.

The practice has been so widespread, especially in Kupat Holim, that its director-general, Asher Yadin, recently said he was willing to pay each department head 50 per cent above his present take-home pay if he would give up his private practice.

"However," said Dr. J. F. Morrissey, Professor of Medicine at the University of Wisconsin, "there was very little interest shown in endoscopy on the part of U.S. pediatricians. We took it up



Lessons Gang Agley

"Something specific to the alcohol molecule" causes addicted mice to forget well-learned lessons, according to Dr. Gerhard Freund (shown with intoxicated mouse), of University of Florida.

Privileges for Top Doctors

To keep outstanding physicians, as well as others, from emigrating from Israel, they were granted various tax benefits, such as car allowances, telephone allowances, and professional literature allowances.

Clomiphene citrate was given on the fifth day of the cycle following either spontaneous or induced bleeding. The initial dose was 50 mg. in one tablet for five days. When ovulation was induced, the drug was not given in the next cycle, and carryover effects were observed.

When ovulation was not induced in the observed cycle, 50 mg. of the agent was given daily for five days after induced bleeding.

When ovulation was not induced in the first cycle, the dosage was increased to 100 mg. daily for five days in the next period.

Co-worker in the study was Dr. Michio Kitagawa.

in adults a little over a year ago and now have what I believe to be the only series so far reported."

On the basis of experience in more than 100 patients, the instrument was found to be preferable for the examination of patients with severe cardiac or pulmonary disease, those who must be examined in bed, and those who are extremely apprehensive or otherwise intolerant to examination with the normal-size endoscope.

Also, Dr. Morrissey said, it has been found useful for following healing in patients with esophagitis, erosive gastritis, and gastric or duodenal ulceration, and for observing effects of drugs in peptic ulcer healing.

The instrument is of special value, he said, in the examination of patients with esophageal or pyloric narrowing

and in adults a little over a year ago and now have what I believe to be the only series so far reported."

On the basis of experience in more than 100 patients, the instrument was found to be preferable for the examination of patients with severe cardiac or pulmonary disease, those who must be examined in bed, and those who are extremely apprehensive or otherwise intolerant to examination with the normal-size endoscope.

Also, Dr. Morrissey said, it has been found useful for following healing in patients with esophagitis, erosive gastritis, and gastric or duodenal ulceration, and for observing effects of drugs in peptic ulcer healing.

The instrument is of special value, he said, in the examination of patients with esophageal or pyloric narrowing

and in adults a little over a year ago and now have what I believe to be the only series so far reported."

On the basis of experience in more than 100 patients, the instrument was found to be preferable for the examination of patients with severe cardiac or pulmonary disease, those who must be examined in bed, and those who are extremely apprehensive or otherwise intolerant to examination with the normal-size endoscope.

Also, Dr. Morrissey said, it has been found useful for following healing in patients with esophagitis, erosive gastritis, and gastric or duodenal ulceration, and for observing effects of drugs in peptic ulcer healing.

The instrument is of special value, he said, in the examination of patients with esophageal or pyloric narrowing

and in adults a little over a year ago and now have what I believe to be the only series so far reported."

On the basis of experience in more than 100 patients, the instrument was found to be preferable for the examination of patients with severe cardiac or pulmonary disease, those who must be examined in bed, and those who are extremely apprehensive or otherwise intolerant to examination with the normal-size endoscope.

Also, Dr. Morrissey said, it has been found useful for following healing in patients with esophagitis, erosive gastritis, and gastric or duodenal ulceration, and for observing effects of drugs in peptic ulcer healing.

The instrument is of special value, he said, in the examination of patients with esophageal or pyloric narrowing

and in adults a little over a year ago and now have what I believe to be the only series so far reported."

On the basis of experience in more than 100 patients, the instrument was found to be preferable for the examination of patients with severe cardiac or pulmonary disease, those who must be examined in bed, and those who are extremely apprehensive or otherwise intolerant to examination with the normal-size endoscope.

Also, Dr. Morrissey said, it has been found useful for following healing in patients with esophagitis, erosive gastritis, and gastric or duodenal ulceration, and for observing effects of drugs in peptic ulcer healing.

The instrument is of special value, he said, in the examination of patients with esophageal or pyloric narrowing

and in adults a little over a year ago and now have what I believe to be the only series so far reported."

On the basis of experience in more than 100 patients, the instrument was found to be preferable for the examination of patients with severe cardiac or pulmonary disease, those who must be examined in bed, and those who are extremely apprehensive or otherwise intolerant to examination with the normal-size endoscope.

Also, Dr. Morrissey said, it has been found useful for following healing in patients with esophagitis, erosive gastritis, and gastric or duodenal ulceration, and for observing effects of drugs in peptic ulcer healing.

The instrument is of special value, he said, in the examination of patients with esophageal or pyloric narrowing

and in adults a little over a year ago and now have what I believe to be the only series so far reported."

On the basis of experience in more than 100 patients, the instrument was found to be preferable for the examination of patients with severe cardiac or pulmonary disease, those who must be examined in bed, and those who are extremely apprehensive or otherwise intolerant to examination with the normal-size endoscope.

Also, Dr. Morrissey said, it has been found useful for following healing in patients with esophagitis, erosive gastritis, and gastric or duodenal ulceration, and for observing effects of drugs in peptic ulcer healing.

The instrument is of special value, he said, in the examination of patients with esophageal or pyloric narrowing

and in adults a little over a year ago and now have what I believe to be the only series so far reported."

On the basis of experience in more than 100 patients, the instrument was found to be preferable for the examination of patients with severe cardiac or pulmonary disease, those who must be examined in bed, and those who are extremely apprehensive or otherwise intolerant to examination with the normal-size endoscope.

Also, Dr. Morrissey said, it has been found useful for following healing in patients with esophagitis, erosive gastritis, and gastric or duodenal ulceration, and for observing effects of drugs in peptic ulcer healing.

The instrument is of special value, he said, in the examination of patients with esophageal or pyloric narrowing

and in adults a little over a year ago and now have what I believe to be the only series so far reported."

On the basis of experience in more than 100 patients, the instrument was found to be preferable for the examination of patients with severe cardiac or pulmonary disease, those who must be examined in bed, and those who are extremely apprehensive or otherwise intolerant to examination with the normal-size endoscope.

Also, Dr. Morrissey said, it has been found useful for following healing in patients with esophagitis, erosive gastritis, and gastric or duodenal ulceration, and for observing effects of drugs in peptic ulcer healing.

The instrument is of special value, he said, in the examination of patients with esophageal or pyloric narrowing

and in adults a little over a year ago and now have what I believe to be the only series so far reported."

On the basis of experience in more than 100 patients, the instrument was found to be preferable for the examination of patients with severe cardiac or pulmonary disease, those who must be examined in bed, and those who are extremely apprehensive or otherwise intolerant to examination with the normal-size endoscope.

Also, Dr. Morrissey said, it has been found useful for following healing in patients with esophagitis, erosive gastritis, and gastric or duodenal ulceration, and for observing effects of drugs in peptic ulcer healing.

The instrument is of special value, he said, in the examination of patients with esophageal or pyloric narrowing

and in adults a little over a year ago and now have what I believe to be the only series so far reported."

On the basis of experience in more than 100 patients, the instrument was found to be preferable for the examination of patients with severe cardiac or pulmonary disease, those who must be examined in bed, and those who are extremely apprehensive or otherwise intolerant to examination with the normal-size endoscope.

Also, Dr. Morrissey said, it has been found useful for following healing in patients with esophagitis, erosive gastritis, and gastric or duodenal ulceration, and for observing effects of drugs in peptic ulcer healing.

The instrument is of special value, he said, in the examination of patients with esophageal or pyloric narrowing

and in adults a little over a year ago and now have what I believe to be the only series so far reported."

On the basis of experience in more than 100 patients, the instrument was found to be preferable for the examination of patients with severe cardiac or pulmonary disease, those who must be examined in bed, and those who are extremely apprehensive or otherwise intolerant to examination with the normal-size endoscope.

Also, Dr. Morrissey said, it has been found useful for following healing in patients with esophagitis, erosive gastritis, and gastric or duodenal ulceration, and for observing effects of drugs in peptic ulcer healing.

The instrument is of special value, he said, in the examination of patients with esophageal or pyloric narrowing

and in adults a little over a year ago and now have what I believe to be the only series so far reported."

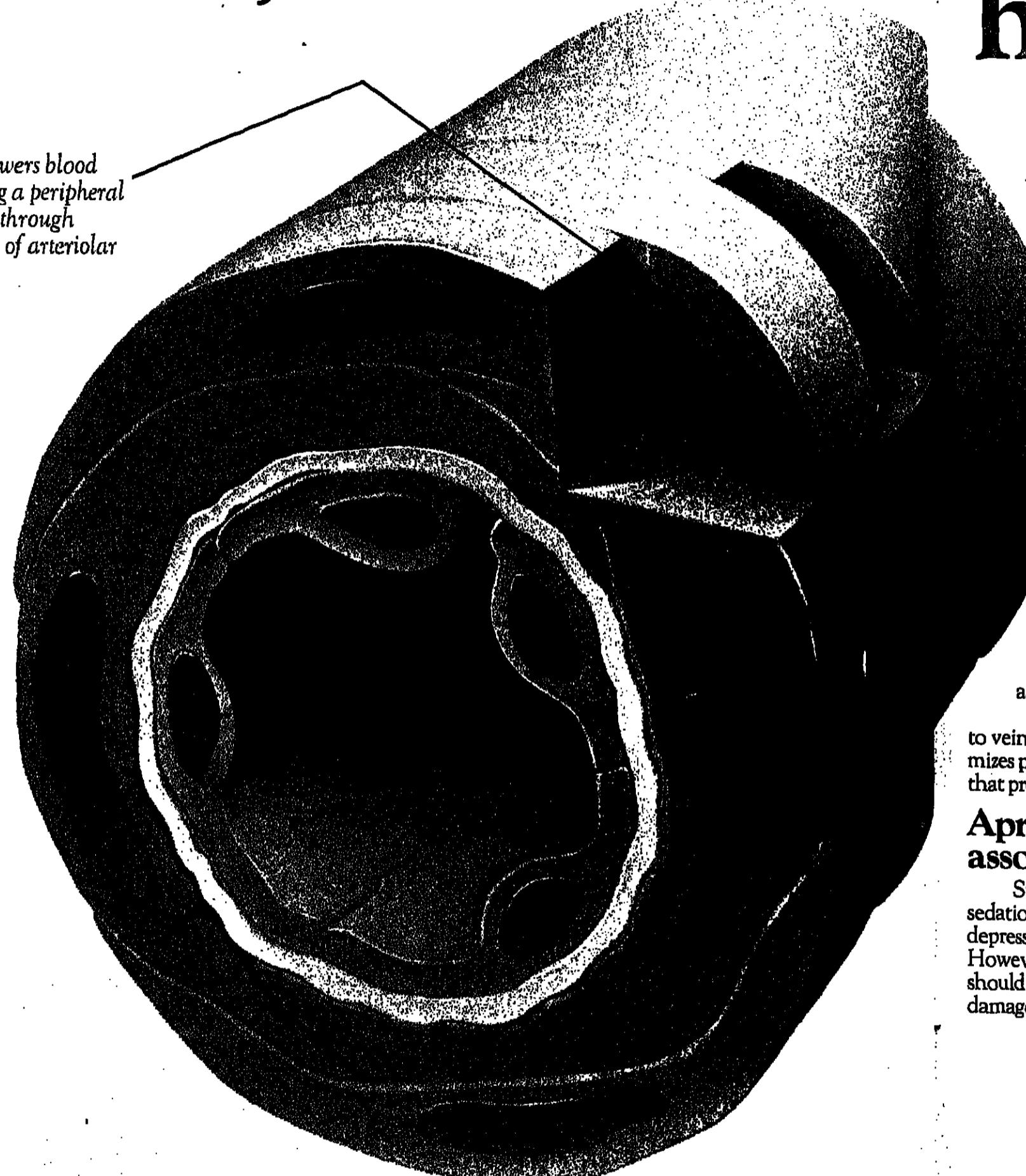
On the basis of experience in more than 100 patients, the instrument was found to be preferable for the examination of patients with severe cardiac or pulmonary disease, those who must be examined in bed, and those who are extremely apprehensive or otherwise intolerant to examination with the normal-size endoscope.

Also, Dr. Morrissey said, it has been found useful for following healing in patients with esophagitis, erosive gastritis, and gastric or duodenal ulceration, and for observing effects of drugs in peptic ulcer healing.

The instrument is of special value, he said, in the examination of patients with esophageal or pyloric narrowing

Apresoline®...where the action is in treating hypertension

Apresoline lowers blood pressure by exerting a peripheral vasodilating effect through a direct relaxation of arteriolar smooth muscle.



An antihypertensive idea whose time has come

Doctors who treat hypertension are increasingly interested in the one oral drug that has a mechanism of action exclusively its own—Apresoline.

Apresoline is in an antihypertensive class by itself because it reduces blood pressure through a unique mechanism. Acting at the ultimate site of hypertension, it directly relaxes arteriolar smooth muscle to decrease peripheral vascular resistance and arterial pressure. As blood pressure falls, there is an accompanying rise in cardiac output and rate.

Apresoline also maintains or increases renal and cerebral blood flow.

Apresoline minimizes postural hypotension

Nickerson¹ describes the action of Apresoline as follows:

"A preferential effect on arterioles, as compared to veins, allows the increase in cardiac output and minimizes postural hypotension; the latter is much less than that produced by agents blocking sympathetic nerves."

Apresoline avoids side effects associated with other agents

Such untoward reactions as drowsiness, lethargy, sedation, sexual dysfunction, and exacerbation of mental depression are not usually encountered with Apresoline. However, as with any antihypertensive agent, hydralazine should be used with caution where advanced renal damage exists.

Apresoline helps tailor the regimen to the patient

When Apresoline is added to an existing antihypertensive regimen, it introduces a different and complementary pharmacologic approach to the control of your patient's hypertension.

Apresoline thus affords the physician a variety of combinations with which he can construct regimens more closely molded to individual requirements. According to Freis,² such a combination of drugs, each with a different antihypertensive mechanism, is the most effective way to control blood pressure. This may also permit lower drug dosages.

Apresoline lends itself admirably to the contemporary antihypertensive rationale and its therapeutic goals: more vigorous and more effective control of blood pressure through a plurality of mechanisms.

Apresoline: used effectively in the VA studies

Apresoline was one of the three basic drugs used in two published VA cooperative studies.^{3,4}

References: 1. Nickerson M: Antihypertensive agents and the drug therapy of hypertension. In Goodman LS, Gilman A (eds): *The Pharmacological Basis of Therapeutics*, ed 4. New York, The Macmillan Company, 1970, p 729. 2. Freis ED: Hypertension: a controllable disease. *Clin Pharmacol Ther* 13:627-632, 1972. 3. Effects of treatment on morbidity in hypertension: Results in patients with diastolic blood pressures averaging 115 through 129 mm Hg. Veterans Administration Cooperative Study Group on Antihypertensive Agents. *JAMA* 202:1028-1034, 1967. 4. Effects of treatment on morbidity in hypertension: II. Results in patients with diastolic blood pressure averaging 90 through 114 mm Hg. Veterans Administration Cooperative Study Group on Antihypertensive Agents. *JAMA* 213:1143-1152, 1970.

Next page: Apresoline (hydralazine) and the Hypertension Task Force

Apresoline® hydrochloride (hydralazine hydrochloride)

TABLETS

INDICATIONS

Essential hypertension, alone or as an adjunct. Hypertension in pulmonary artery disease; mitral valve and rheumatic heart disease.

WARNINGS

Chronic administration of doses over 500 mg per day may produce an arteriolar syndrome leading to a clinical picture simulating acute systemic lupus erythematosus. This may also occur at lower doses. Most of these reactions are reversible upon withdrawal of therapy, but long-term treatment with hydralazine may be necessary and should have been discontinued if the following complete blood counts, L.E. cell preparations and serum nuclear antibody titer determinations are indicated periodically during prolonged therapy, even though they are asymptomatic. These studies are also indicated in the presence of any unexplained symptoms.

Use MAO inhibitors with caution.

Usage in Pregnancy
The drug should be used only when, in the judgment of the physician, it is deemed essential to the patient.

PRECAUTIONS

Use cautiously in suspected coronary artery or other cardiovascular diseases, cerebral vascular disease, and renal damage. Postural hypertension may occur. The pressor response to epinephrine may be reduced. Peripheral neuritis, evidenced by paresthesias, numbness, and tingling, has been observed. Published evidence suggests an antipyridoxine effect

and addition of pyridoxine to the regimen if symptoms develop.

Blood dyscrasias, consisting of reduction in hemoglobin and red cell count, leukopenia, agranulocytosis, and purpura, have been reported.

If such reactions occur, discontinue the drug. Periodic blood counts are advised during prolonged therapy.

ADVERSE REACTIONS

Common adverse reactions: palpitations, anorexia, nausea, vomiting, diarrhea, tachycardia, orthostatic hypotension, constipation, epigastric pain, headache, dizziness, tinnitus, and rarely, hepatitis; constipation; difficulty in micturition; transient paralytic lesions; lymphadenopathy; splenomegaly; peripheral neuritis; alopecia, consisting of reduction in hemoglobin and red cell count, leukopenia, agranulocytosis, and purpura; hypertension; paroxysmal pressor responses.

evidenced by paresthesias, numbness, and tingling; headache; dizziness; tremors; muscle cramps; depression, or anxiety; hypertension (including rash, urticaria, pruritus, fever, chills, arthralgia, and rarely, hepatitis); constipation; difficulty in micturition; transient paralytic lesions; lymphadenopathy; splenomegaly; peripheral neuritis; alopecia, consisting of reduction in hemoglobin and red cell count, leukopenia, agranulocytosis, and purpura; hypertension; paroxysmal pressor responses.

DOSAGE
Initiate therapy in gradually increasing dosages, adjust according to individual responses. Start with 25 mg 4 times daily for the first week. Increase to 25 mg 4 times daily for balance of first week. For second and subsequent weeks, increase dosage to 50 mg 4 times daily. For maintenance, adjust dosage to individual needs.

The incidence of toxic reactions, particularly the L.E. cell syndrome, is high in the group of patients receiving large doses of Apresoline. In such cases, a lower dosage of Apresoline combined with a thiazide, reserpine, or both may be considered. However, when combination therapy is used, the physician should insure the lowest possible therapeutic dose of each drug.

HOW SUPPLIED

Tablets, 100 mg (peach, dry-coated); bottles of 100. Consult complete literature before prescribing.

CIBA Pharmaceutical Company
Division of CIBA-GEIGY Corporation
Summit, New Jersey 07901

C I B A

Apresoline... (hydralazine)

part of the Hypertension Task Force "plan of action"

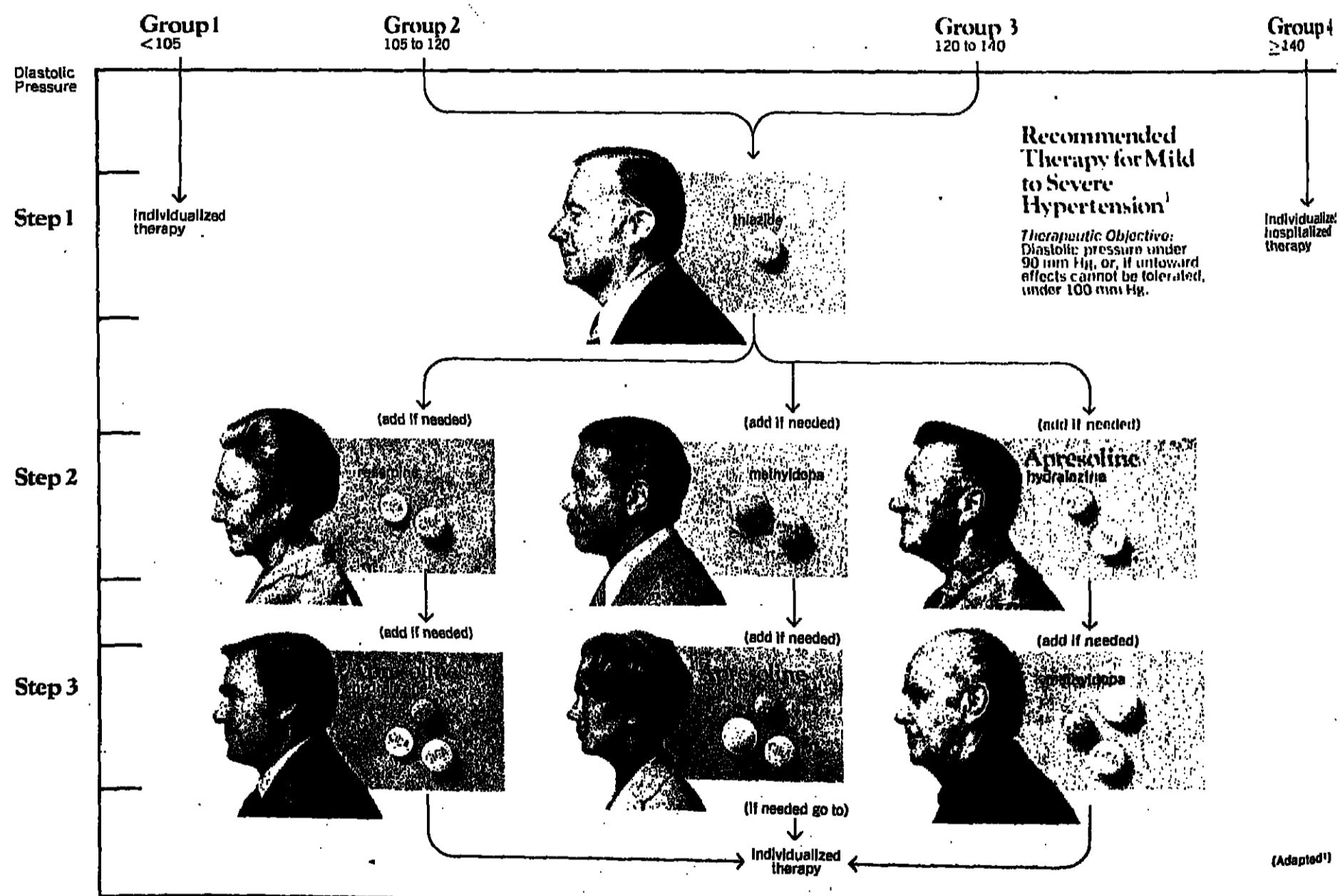
In September 1973, Task Force I of the National High Blood Pressure Education Program recommended a series of antihypertensive regimens for groups with hypertension ranging from mild to severe. Hydralazine—used in combination with sympathetic-inhibiting and/or diuretic antihypertensive

agents—was a specific recommendation for "second step" and "third step" therapy in patients with diastolic pressures ranging from 105 to 140 mm Hg.

Hydralazine played a prominent role in the Task Force regimens' because of its compatibility with almost any antihypertensive regimen. For

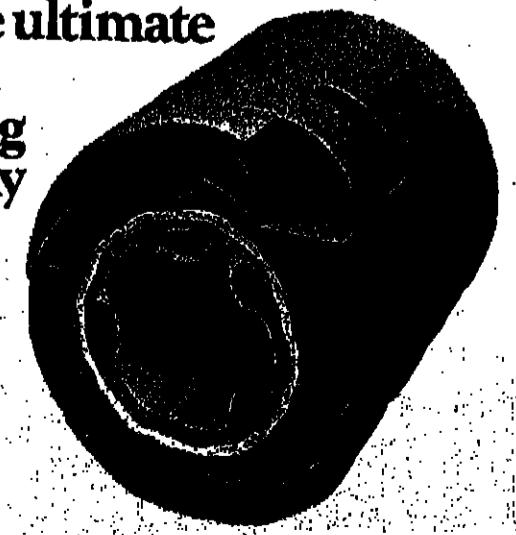
Apresoline can be combined advantageously with nearly all diuretics and sympathetic inhibitors.

Reference: 1. Report of Task Force I, National High Blood Pressure Education Program: Recommendations for a National High Blood Pressure Program Data Base for Effective Antihypertensive Therapy, Sept 1, 1973. DHEW Publication No. (NIH) 74-535.



Apresoline® (hydralazine)
...acts directly at the ultimate
site of hypertension
...brings something
special to almost any
antihypertensive
regimen

For brief prescribing information,
please see preceding page.



C I B A

Wednesday, February 5, 1975

MEDICAL TRIBUNE

Check, Double Check Breast Ritual 'Still Best'

Medical Tribune Report

BOSTON—A combination of self-examination and twice-a-year checkups by a physician is still the most effective method for the early detection of breast cancer, Dr. Richard Wilson told a Harvard Medical Society symposium here.

"I'm afraid this somewhat quaint ritual is here to stay until we have a no-fail test, such as a blood test," he said.

Dr. Wilson, who is Associate Professor of Surgery at the Peter Bent Brigham Hospital, pointed out that although xeroradiography and thermography are effective for early diagnosis in patients who are at risk because of their age, they have not proved their worth when used for the younger woman.

"There is a great danger today to put too much faith in these techniques," Dr. Wilson warned the audience of students and physicians.

He reminded them that there is a great deal of fibrocystic disease in most breasts and that the breasts change constantly through the monthly cycle.

"The real job is to decide that what you detect is a matter for concern," he remarked.

More Aspirations In Office

Dr. Wilson said that he is doing "more cystic aspirations in my office than ever before; otherwise I biopsy all mass lesions—regardless of what the screening says."

Dr. Lester Kalisher said that while xeroradiography can reveal a cancerous or precancerous lesion before it becomes palpable, the barely palpable 2-cm. mass today is considered a late symptom.

At the Massachusetts General Hospital, where he is an Instructor in Radiology, xeroradiography is used in women who present symptoms or are considered to be at high risk because of family history, age, or earlier lesions.

"What we look for are the microcalcifications without mass," Dr. Kalisher said. "Eighty per cent of these malignancies have such calcification."

Physicians at M.G.H., he added, also look for asymmetric duct patterns—unusual duct outlines that appear on one side of the breast and not on the other, and are easy to spot by xeroradiography because both sides are presented at the same time.

Of the 1,315 referrals for xeroradiography made at the hospital so far, he reported, 125 were recommended

Next In Consultation

DR. CHARLES M. PLOTZ, Chairman, Department of Family Practice and Director of Continuing Education, Professor of Medicine, State University of New York Downstate Medical Center, Brooklyn, N.Y., . . . will answer questions on what's new and important in diagnosing and treating polymyalgia rheumatica and rheumatoid arthritis, diagnostic criteria, whether a temporal artery biopsy should be done in diagnosis of polymyalgia rheumatica, length of treatment and sequence of treatment.

for biopsy. Sixty-four of the lesions proved malignant, 33 benign, and the rest were not biopsied.

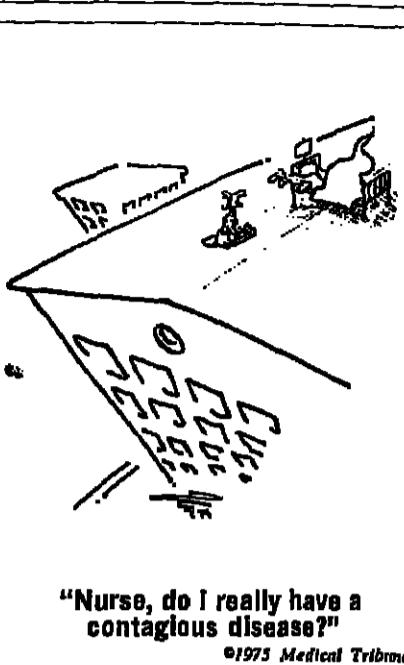
Dr. Norman L. Sadowsky, radiologist in chief at the Faulkner Hospital, said that thermography is the preferred diagnostic tool at his institution.

Thermography picks up some carcinomas that xeroradiography does not, he said, and further, the method is more practical for annual examinations. It takes little time—about 10 minutes—and is so inexpensive to use that Faulkner does not charge for it.

Charges for a xeroradiography examination in Boston, it was noted, range from \$50 to \$100.

Initial costs for installation also differ considerably, although in the other direction, the seminar was told. The

In the past three years nearly half of all women over 30 have been examined, resulting in four times as many diagnoses of cancer as in the previous period and eight times as many patients identified with cancer in its very early stages.



"Nurse, do I really have a contagious disease?"

©1975 Medical Tribune

Clinical supply available on your request

Hycomine® Syrup

Each teaspoonful (5 ml) of orange-colored, fruit-flavored syrup contains hydrocodone bitartrate (Warning: May be habit forming) 5.0 mg, homatropine methylbromide 1.5 mg, pyramine maleate 2.5 mg, phenylephrine hydrochloride 10.0 mg, ammonium chloride 60.0 mg.

Hycomine® Rx's may be
refilled five times within 6 months[†]

Telephone prescriptions
permitted in most states[†]



To receive your clinical supply, please fill out this coupon (Being sure to include your BNDD number and signature) and mail to:

Professional Request Dept.
Endo Laboratories, 1000 Stewart Avenue
Garden City, NY 11530

Name (please print)

Street

City

BNDD Number

Physician Signature

Your name, address and BNDD Number must be legible and complete for us to honor your request.

Where permitted by state laws and regulations.

†Where permitted by state laws and regulations.

Endo Laboratories, Inc.
Subsidiary of E.I. du Pont de Nemours & Co., Inc.
Garden City, New York 11530

Sex Hormone Combo Proposed for Male Pill

Medical Tribune World Service

MELBOURNE—Dr. Michael Briggs, director of biochemistry at the Alfred Hospital, and Dr. Maxine Briggs, assistant medical superintendent, said that a combination estrogen-androgen could be the answer to the search for a safe and effective male contraceptive pill free of the side effects of loss of libido and testicular atrophy.

Dr. Michael Briggs related that he became interested in this approach when he discovered that two elderly osteoporosis patients who were taking an estrogen-androgen combination developed severely reduced sperm production.

Then, five healthy volunteers were selected and a trial started with two

pills being taken daily at meals. By day 63 of hormone treatment, four of the five men had become infertile, and during the 18th week, the fifth patient also became infertile.

No Pregnancies in 16 Weeks

The treatment was maintained for 34 weeks and the volunteers' wives went off their oral contraceptives from week 18. No pregnancies resulted in the 16 trial weeks. Sperm production was back to normal within five weeks after discontinuation of hormone treatment.

As a control for questions on sex drive, the volunteers were given placebo tablets for three weeks at the start of the course.

Two volunteers reported decreased libido in the first eight weeks, which included the placebo period, but normal libido for the remainder of the study.

Another man reported increased libido during the second half of the treatment period, while he and another subject experienced a reduction in libido after treatment was stopped.

Three men reported occasional mild nausea while they were taking the pill.

Dr. Briggs said there were no changes in skin, hair, breasts, or urination.

With further refinement, he said, the pill could be developed to be taken less frequently—every other day or a few times every two weeks.

Now, for both aspects of constipation



Announcing

Senokot® S Tablets

(standardized senna concentrate and dioctyl sodium sulfosuccinate)

a unique natural laxative plus
a classic stool softener

Provides a unique natural laxative—standardized senna concentrate... virtually colon-specific... effectiveness documented in numerous published studies comprising thousands of patients.

Provides a classic stool softener—DSS... complementing the laxative action by softening the stool for smoother and easier passage.

Comfortable, predictable evacuation... a bedtime dose of SENOKOT S Tablets usually induces comfortable evacuation the next morning, allowing uninterrupted sleep. SENOKOT S Tablets aid in rehabilitation of the constipated patient by facilitating regular elimination.

Indications: SENOKOT S Tablets offer welcome relief in functional constipation when combined neuromotor stimulation plus stool softening is indicated, especially for the aged, postpartum and postoperative patients; drug-induced constipation; constipated patients and those with hemorrhoids. Dose: (preferably bedtime): Adults: 1 tablet max. Children: 1/2 tablet. Individual requirements, dosage may be decreased. **Contraindications:** Do not use in patients with the most sensitive bowel. **Warnings:** Do not use in patients with intestinal obstruction, rectal bleeding, and intestinal perforation. **Precautions:** Do not use in patients with intestinal obstruction, rectal bleeding, and intestinal perforation. **Side Effects:** Do not use in patients with intestinal obstruction, rectal bleeding, and intestinal perforation. **Adverse Reactions:** Do not use in patients with intestinal obstruction, rectal bleeding, and intestinal perforation. **Storage:** Do not use in patients with intestinal obstruction, rectal bleeding, and intestinal perforation. **Manufactured by:** The Procter & Gamble Company, Cincinnati, Ohio 45201. © 1974 The Procter & Gamble Company.

MEDICAL AUDIOVISUAL SERVICES

... brief summaries of editorials or comments in current medical and scientific journals.

Disaster Management

"... if a fully loaded aircraft flying over a city should crash on a residential area or in the city centre, it could produce a casualty list approaching five thousand. Such a disaster would require the help of the armed forces and their medical services. Thought must also be given to the possibility of nuclear disaster. This is again an area where there is very little experience, but every hospital should have some idea how it would cope with decontamination after nuclear fall-out—largely a matter of providing a special area with a plentiful supply of water." (Special article, *David Cope; The Lancet* 2:1309, Nov. 30, 1974)

Delay in Energy Sources

"Consumption of energy goes on unabated in spite of a recession, higher prices, and presidential appeals. But domestic reserves of hydrocarbons are being depleted rapidly and the stage is being set for empty gasoline pumps, cold homes, and large-scale unemployment unless there is a drastic change in attitudes soon. A major factor is the long time span involved in creating new sources of energy. . . .

"The first reactor went critical in December 1942. In 1973, nuclear energy accounted for only 1 percent of the nation's energy consumption. Ten years from now, nuclear energy will meet at most 7 percent of the nation's needs. . . .

"Thus, for at least the next decade, energy horizons will be limited by oil, natural gas, and coal. But available domestic supplies of oil and gas are diminishing, at the rate of 4 to 6 percent per year for oil and about 7 to 8 percent per year for natural gas. . . .

"Perhaps the most serious and certainly the least recognized problems lie in the supplies of natural gas. It heats 55 percent of the nation's homes, is widely used as a feedstock for petrochemicals, including fertilizer, and is by far the largest source of energy for industry . . . equivalent to that of about 5 million barrels of oil. National policy accords priority to residential demand for natural gas. The rate of decay of supplies is such that by 1980, with a few exceptions, industry will be prevented from using natural gas. This would have enormous effects on the economy.

"To make good the energy deficit due to decay of natural gas alone, a doubling of coal production during the next 6 years would be required. But to open a new underground mine requires about 5 years. The quickest path toward relief is expansion of surface mining of low-sulfur coal in the Rocky Mountain States. But with various delays connected with changes from gas or oil to coal and with environmental controversies, heaven only knows when this country will emerge from the years of travail; and discontent that it is now entering." (Editorial, *Philip H. Abelson; Science* 187:17, Jan. 10, 1975)

Blunt Chest Trauma Cited As Cause of Pneumatoceles

Medical Tribune Report

GALVESTON, TEX.—Traumatic lung and paramediastinal pneumatoceles are "not well appreciated" as a manifestation of nonpenetrating chest trauma, according to a radiologist at the University of Texas Medical Branch.

These lesions are "definite, acutely formed, primary structural manifestations" of injury, and not secondary lesions—that is, they do not result from the resolution of a pulmonary hematoma—according to Dr. Charles J. Fagan, Associate Professor of Radiology.

"Awareness of this fact," he said, "will explain the (common) finding of a cyst, often containing an air-fluid level, on the initial or emergency room roentgenogram" of patients who have suffered nonpenetrating chest trauma, most commonly from an automobile accident.

Hemoptysis frequently follows the accident and may be seen during the initial physical examination of the patient, he noted.

In general, Dr. Fagan said, patients are asymptomatic, and the pneumatoceles eventually disappear with no treatment; they last from about one week to as long as three months.

The roentgenographic appearance of the traumatic pneumatocele varies according to its location, whether it holds blood, and whether it is associated with a pulmonary contusion, he observed.

It may be completely opacified and appear as a solitary pulmonary nodule, like that of a hematoma, he said, but

New Japanese Audiovisual Teaching Aid



A Japanese medical student uses one of the new audiovisual units for teaching x-ray diagnosis developed by the Japan Research Center. She pushes a button to turn on a physician discussion of the x-ray she is studying.

NEW
from Wyeth

wygesic

each tablet
contains

65 mg. propoxyphene HCl, U.S.P.
and
650 mg. acetaminophen, N.F.

Wyeth

© 1975 The Procter & Gamble Company

Induced Abortions Reported To Boost Risk of Spontaneous

Medical Tribune World Service

PRAGUE — Artificial termination of pregnancy greatly increases the risk of a subsequent spontaneous abortion, Prof. Alfred Kotásek, head of the Gynecological and Obstetrical Clinic of Charles University, Prague, told the Fourth European Congress of Perinatal Medicine here.

It also enhances the likelihood of premature births and ectopic pregnancies, he said. Further, "abortion frequently reduces woman's future reproductive capability and affects her emotional and sexual life."

He warned that in a review of the literature, "a great sum of serious morbidity following legal artificial termination of pregnancy has been noticed and described in many papers." Most clinics, he said, lose sight of their patients soon after the operation, but long-term studies including subsequent pregnancies are necessary for a true picture of post-abortion complications.

2 MILLION Abortions in 17 Years

Czechoslovak experience is based on some 2,000,000 legal first trimester abortions (voluntary induced abortions are not permitted after the twelfth week), carried out over a period of 17 years. During the first ten years there were 20 maternal deaths connected with the procedure, Dr. Kotásek said, (two per 100,000); since then the rate has decreased.

However, he said, a detailed Prague study concludes that only 57 per cent of pregnancies following induced abortion were carried to term. The spontaneous abortion rate was 2.2 times the "normal" incidence. While reports of cervical incompetence was a rare cause of second trimester miscarriages before legalization of abortion in Czechoslovakia in 1958, ten years later it was reported two to five times more frequently in women who had had interruptions than those who had not. "A very high standard of antenatal care from the end of the first trimester for all women who have had a previous artificial termination of pregnancy is advisable," he said.

Czechs Smoking More

Medical Tribune World Service

PRAGUE—Despite a policy of no tobacco advertising, despite antismoking clinics, and despite the publicity given to the harmful effects of the weed, cigarette smoking continues to increase in Czechoslovakia. Cigarette sales have tripled since 1946 and now amount to 27 billion annually, or 1900 per capita.

Much of the increase is accounted for by women and children. According to an investigation recently published by the Institute of Health Education in Prague, boys now try their first cigarette before they are ten, girls between the ages of twelve and 13. By the time they are fifteen, every second youngster has at least tried smoking, and every fourth smokes occasionally.

Officials attribute 30,000-40,000 deaths a year, about a fifth of all deaths, to smoking-connected causes.

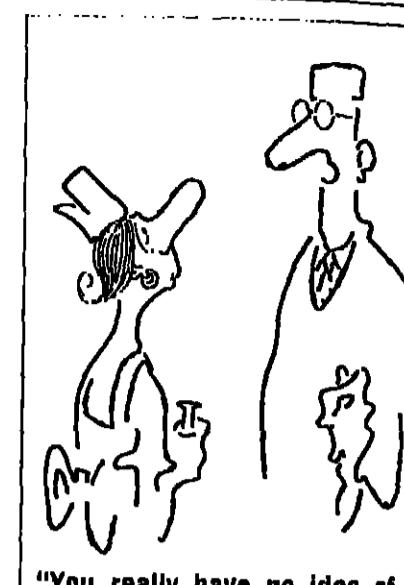
abortion, since women who do not wish children do not attend fertility clinics, Czechoslovak authors report a sterility rate of 1.3-7 per cent following induced abortion, compared with 2-5 per cent reported elsewhere.

Sex Attitudes Changed in 30%

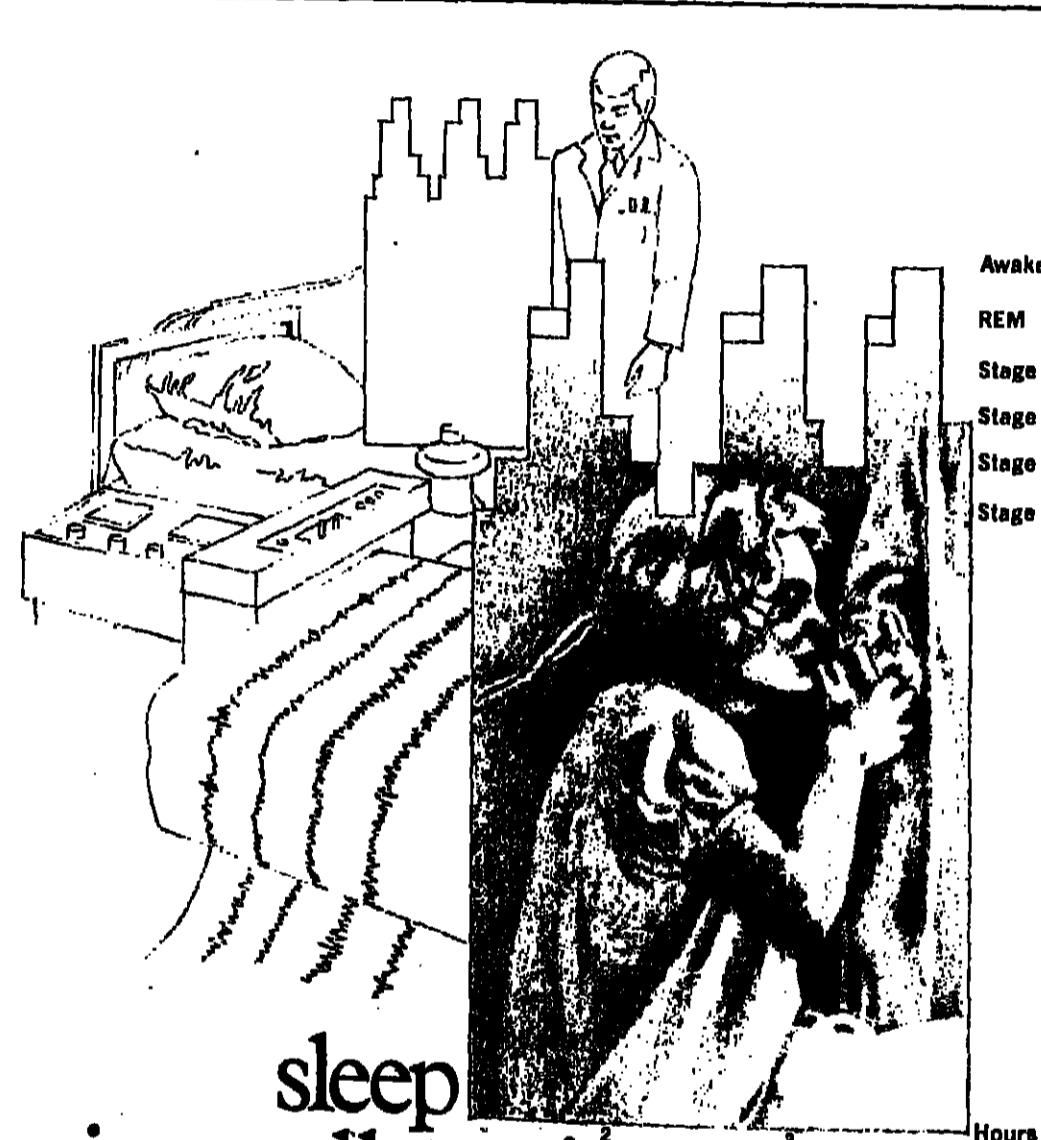
Functional sexual disorders are also a common late consequence. Of 200 women who were examined psychologically by one Czechoslovak author, in connection with interruption of pregnancy, more than 30 per cent admitted lower or negative attitudes towards sexuality.

Significant increases in the duration of the third stage of labor and in retained and adherent placenta have also been reported in women who had previously had induced abortions, Dr. Kotásek said.

Although it is impossible to obtain complete figures on sterility following



"You really have no idea of the difference between fallopian tubes and ureters?"
©1975 Medical Tribune



sleep
Hours
is usually maintained with
fewer nighttime awakenings...
a consistent benefit of

Dalmane
(flurazepam HCl) proved by a
17-night clinical study in the sleep research
laboratory evaluating effectiveness in
insomnia patients!

Eight patients received no medication on nights 1-4; Dalmane (flurazepam HCl) or placebo on nights 5-9; crossover capsule, nights 10-14; and no medication, nights 15-17. While placebo had no significant effect on sleep maintenance, Dalmane reduced nighttime awakenings by 55.1% when given on nights 5-9, 43.7% on nights 10-14. When four control subjects received placebo on the 10 "drug" nights, awakenings increased 11.5% over baseline.

Wednesday, February 5, 1975

MEDICAL TRIBUNE

IFRP Intrauterine Membrane Disappointing

Medical Tribune World Service

5.9 per 100 women, the pregnancy rate was 1.1, and the expulsion rate was 2.0.

The pleated-membrane IUD is a polyethylene device containing 15 per cent barium sulfate. It is approximately 1.5 inches long and 0.005 inch thick. The pleats were designed to increase the device's ability to react to uterine contractions. The IUD is strengthened by a "wishbone" reinforcement molded on the bottom.

About half of the study group were less than 25 years old and about 80 per cent had one or two children.

Fourteen of the 119 women tested had the IUD removed because of bleeding, said Dr. Thomas, research assistant with the IFRP of the Carolina Population Center, University of North Carolina.

After three months, the net cumulative rate of bleeding/pain removals was

not only to develop an improved IUM," said Dr. Thomas, "but also to develop hypotheses concerning the mechanism of action which leads to increased or decreased bleeding in all IUDs."

Coauthors were Drs. Leonard Laufe, of the Western Pennsylvania Hospital, Pittsburgh, and Robert Wheeler, of the Battelle Memorial Institute, Richland, Wash.

Latex-Leaf IUD

► Israeli doctors, on the basis of initial results, pronounced the Anderson-Ansell latex-leaf IUD superior to the Lippes loop and Dalkon shield in some respects—notably in low pregnancy rates. Dr. E. Sadovky, of the Hadassah University Hospital, Jerusalem, re-

ported on 187 women from 18 to 40 years in whom the latex leaf was inserted between January, 1973, and March, 1974, and who used it for a total of 1,712 women-months.

The latex leaf IUD is made of inert silicone rubber impregnated with copper and zinc and is radiopaque, Dr. Sadovky said. The electromechanical interaction of the metallic ion it releases is believed to cause its contraceptive effect.

Its softness was expected to prevent decubitus and irritation of the uterus, with consequent low removal rates due to bleeding and pain. But this did not prove to be the case.

High Removal Rate

The removal rate was 37.1 per 100 woman-years, compared with 28.9 for the Lippes loop and 14.1 for the Dalkon shield, Dr. Sadovky reported. Removal was mainly due to bleeding—23.1 with the latex leaf, against 12.8 with the loop and 6.4 with the shield.

The pregnancy rate, however, was only 1.4, against 12.3 and 4.2 respectively with the two other devices.

The expulsion rate was 4.2 against 12.3 and 1.43.

The investigators commented that the low pregnancy rate, the ease of insertion, and the fact that in some patients with high parity and with slightly enlarged uterus there is relatively little side-effect bleeding, make it worthwhile to try the latex-leaf IUD in larger groups of women.

Coauthors were Drs. W. Z. Polishuk, S. O. Antebi, S. Yarkoni, and Y. Aboulafia.

No Ideal Topical Drug Seen for Tinea Pedis

Medical Tribune Report

CHICAGO—Patients who present with symptoms of athlete's foot are best treated by basic hygienic measures and steps to keep the feet cooler, such as loose shoes, sandals, or leaving the shoes off frequently, according to Dr. Leon Goldman, Professor and Chairman of the Department of Dermatology, University of Cincinnati.

"In spite of extensive advertisements in the lay press and television, there is still no ideal type of topical medication for athlete's foot," he told the American Academy of Dermatology.

Renewed attention is being given to topical griseofulvin. With suitable vehicles, the medication may have some value, but further control studies are needed, Dr. Goldman said. Newer synthetic agents available to the practitioner include halopropen, miconazole nitrate and silver sulfadiazine.

Topical medications should be continued for some time after symptoms improve, unless they are irritating or sensitizing, Dr. Goldman recommended.

Preventive measures should include "simple drying of the skin without using the towel as a saw to tear the skin between the toes," he added. Bland powders are helpful.

He pointed out that the combination of poor hygiene through heavy, sweaty socks, especially nylon and wool, and heavy shoes provides favorable moist conditions for the continued growth of the fungus infection.

confirmed by clinical studies in four geographically separated sleep research laboratories^{2,3}

Using a 14-night protocol, involving eight insomniacs and eight normal subjects, four studies confirmed the sleep-maintaining effectiveness of Dalmane (flurazepam HCl) and the reproducibility of this response. On average, one 30-mg capsule reduced number of awakenings by 31.3% and wake time by 52.6%. In all these studies, Dalmane induced sleep rapidly, on average within 17 minutes; reduced nighttime awakenings; and provided, on average, 7 to 8 hours of sleep without repeating dosage.^{2,3}

Dalmane (flurazepam HCl) induces and maintains sleep, with relative safety

Dalmane is generally well tolerated; morning "hung-over" has been relatively infrequent. While dizziness, drowsiness, lightheadedness and the like have been noted most often, particularly in the elderly and debilitated, physicians should be aware of the possibility of more serious reactions, as noted in the Complete Product Information.

Before prescribing Dalmane (flurazepam HCl), please consult Complete Product Information, a summary of which follows:

Indications: Effective in all types of insomnia characterized by difficulty in falling asleep, frequent nocturnal awakenings and/or early morning awakening; in patients with recurring insomnia or poor sleeping habits; and in acute or chronic medical situations requiring restful sleep. Since insomnia is often transient and intermittent, prolonged administration is generally not necessary or recommended.

Contraindications: Known hypersensitivity to flurazepam HCl.

Warnings: Caution patients about possible combined effects with alcohol and other CNS depressants. Caution against hazardous occupations requiring complete mental alertness (e.g., operating machinery, driving). Use in women who are or may become pregnant only when potential benefits have been weighed against possible hazards. Not recommended for use in persons under 15 years of age. Though physical and psychological dependence have not been reported on recommended doses, use caution in administering to addiction-prone individuals or those who might increase dosage.

Precautions: In elderly and debilitated, initial dosage should be limited to 15 mg to preclude oversedation, dizziness and/or ataxia. If combined with other drugs having hypnotic or CNS-depressant effects, consider potential additive effects. Employ usual precautions in patients who are severely depressed, or with latent depression or suicidal tendencies. Periodic blood counts and liver and kidney function tests are advised during repeated therapy. Observe usual precautions in presence of impaired renal and/or hepatic function.

Adverse Reactions: Dizziness, drowsiness, lightheadedness, staggering, ataxia and falling have occurred, particularly in elderly or debilitated patients. Severe sedation, lethargy, disorientation and coma, probably indicative of drug intolerance or overdose, have been reported. Also reported were headache, heartburn, upset stomach, nausea, vomiting, diarrhea, constipation, GI pain, nervousness, talkativeness, apprehension, irritability, weakness, palpitations, chest pains, body and joint pains and GU complaints. There have also been rare occurrences of sweating, flushes, difficulty in focusing, blurred vision, burning eyes, faintness, hypotension, shortness of breath, pruritus, skin rash, dry mouth, bitter taste, excessive salivation, anorexia, euphoria, depression, slurred speech, confusion, restlessness, hallucinations, and elevated SGOT, SGPT, total and direct bilirubins and alkaline phosphatase. Paradoxical reactions, e.g., excitement, stimulation and hyperactivity, have also been reported in rare instances.

Dosage: Individualize for maximum beneficial effect. **Adults:** 30 mg usual dosage; 15 mg may suffice in some patients. **Elderly or debilitated patients:** 15 mg initially until response is determined. **Supplied:** Capsules containing 15 mg or 30 mg flurazepam HCl.

One 30-mg capsule h.s. — usual adult dosage (15 mg may suffice in some patients).

One 15-mg capsule h.s. — initial dosage for elderly or debilitated patients.

when restful sleep is indicated

Dalmane

(flurazepam HCl)

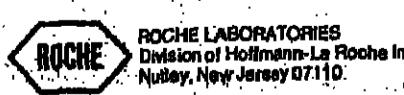
One 30-mg capsule h.s. — usual adult dosage (15 mg may suffice in some patients).

One 15-mg capsule h.s. — initial dosage for elderly or debilitated patients.

• induces sleep within 17 minutes, on average

• reduces nighttime awakenings

• sustains sleep 7 to 8 hours, on average, without repeating dosage



2. Kales J, et al: *Clin Pharmacol Ther* 12:691-697, Jul-Aug, 1971

3. Karcan I, Williams RL, Smith JR: The sleep laboratory in the investigation of sleep and sleep disturbances, Scientific exhibit at the 124th annual meeting of the American Psychiatric Association, Washington DC, May 3-7, 1971

4. Frost JD Jr: Data on file, Medical Department, Hoffmann-La Roche Inc, Nutley NJ

5. Dement WC: Data on file, Medical Department, Hoffmann-La Roche Inc, Nutley NJ

ROCHE LABORATORIES
Division of Hoffmann-La Roche Inc.
Nutley, New Jersey 07110

Mexicans Describe Fatal Muscle Hypertonia

Medical Tribune World Service

MEXICO CITY—An unusual neurologic disorder, previously undescribed, consisting of severe generalized muscle hypertonia during wakefulness and normotonia during sleep, has been reported by Mexican investigators.

Drs. Jose Maria Cantu, of the genetics section, biology of reproduction division, and Alfredo Cuellar, head of the department of nutrition, Hospital de Pediatría, Centro Medico Nacional, Mexican Social Security Institute, described one case in which the condition manifested itself at birth and remained unchanged until the infant's death two and one-half months later from bronchopneumonia.

The body stiffness was such that the patient could be moved from dorsal

decubitus to an erect position by supporting him only by his feet and at the nape of the neck. The arms were in flexion, the hands strongly clenched, and the feet in hyperflexion. He remained in that state all the time he was awake; after falling asleep, he gradually relaxed.

6 Sibs Affected

Dr. Cantu concluded from the family study that homozygosity of a mutant recessive gene located in an autosome was responsible for the disease. Six sibs of both sexes were indirectly ascertained to have been similarly affected and to have died of the disorder between two and four month of age. The parents were second cousins and had 19 other children.

Attempts to correct the hypertonia with intravenous administration of calcium gluconate two days after birth had no effect, nor did methocarbamol intramuscularly at two months of age, but a week later, a single dose of benzodiazepine produced a mild reaction.

The neuromuscular impairment had resulted in fetal hypokinesis. After the birth of the second affected child, Dr. Cantu said, the mother was able to predict which of the subsequent children would be likewise affected on the basis of the weak fetal movements she felt.

Also present in the infant studied were pharyngoesophageal dyskinesia and cardiopulmonary distress, complicated by bronchopneumonia unresponsive to treatment.

Merrell

Tenuate® (diethylpropion hydrochloride N.F.)

BRIEF SUMMARY
INDICATIONS: Tenuate is indicated in the management of exogenous obesity as a short-term adjunct (a few weeks) in the treatment of weight reduction based on caloric restriction. The first few weeks of treatment, caloric intake should be measured against possible caloric inherent in their diet.
CONTRAINDICATIONS: Advanced arteriosclerosis, hypertension, known hypersensitivity, or idiosyncrasy to the sympathomimetic drugs, glucose, Aspirated states.

Patients with a history of drug abuse. During the 14 days following the administration of monoamine oxidase inhibitors, hypertensive crises may result.

WARNING: If tolerance develops, the recommended dose should not be exceeded in an attempt to increase the effect; tolerance may be induced. Tenuate may impair the ability of the patient to perform potentially hazardous activities such as operating machinery or driving an automobile; the patient should therefore be cautioned accordingly.

Drug Dependence: Tenuate has some chemical and pharmacologic similarities to the amphetamines and other related stimulants. Drugs that have been extensively abused. There may be an increased risk of dependence on amphetamines later chronically abusive drugs. The possibility of abuse should be kept in mind when evaluating the results of Tenuate. The amphetamines and related drugs may be associated with varying degrees of psychologic dependence and social dysfunction which is the case with all drugs, may be severe. There are reports of patients who have inappropriately come to rely on these recommended. Abrupt cessation following high dosage administration results in extreme fatigue and mental confusion; changes are also noted on the sleep EEG. Maculopapular rash and exfoliation with anesthetic drugs include severe dermatitis, marked irritability, hyperactivity, and personality changes. The most common manifestation of chronic intoxication is psychosis, often clinically indistinguishable from schizophrenia.

Use in Pregnancy: Although rat and human reproductive studies have not indicated adverse effects, the use of Tenuate during pregnancy or may become pregnant requires that the potential benefits be weighed against the potential risks.

Use in Children: Tenuate is not recommended for use in children under 12 years of age.

PRECAUTIONS: Tenuate should be exercised in prescribing Tenuate for patients with hypertension or with symptomatic cardiovascular disease, including arrhythmias. Tenuate should not be administered to patients with severe hypertension, hypotension, or hypotension. Tenuate insulin requirements in diabetes mellitus may be altered in association with the use of Tenuate and the concomitant dietary regimen.

Tenuate may decrease the hypotensive effect of guanethidine. The dose of guanethidine should be decreased or dispensed at one time in order to minimize the possibility of over dosage. Reports suggest that Tenuate may increase convulsions in some epileptics. Therefore, epileptics receiving Tenuate should be closely monitored. Titration of dose or discontinuance of Tenuate should be gradual.

ADVERSE REACTIONS: Cardiovascular: Palpitation, tachycardia, elevation of blood pressure, precardial pain.

Central Nervous System: Overstimulation, nervousness, restlessness, dizziness, tinnitus, insomnia, anxiety, euphoria, depression, tremor, tics, headache; rarely psychoses episodes of hallucinations delusions, few epileptics an increase in convulsive episodes has been reported.

Gastrointestinal: Dryness of the mouth, constipation, constipation, other gastrointestinal disturbances.

Adverse: Urinary, rash, eczema, asthma.

Endocrine: Impotence, changes in libido, menstrual cramps.

Hematologic: Bone marrow depression, agranulocytosis, leukopenia.

Metabolic: A variety of miscellaneous adverse reactions have been reported by physicians. These include disorders such as dyspepsia, hair loss, muscle pain, dysuria, and polyuria.

MERRELL-NATIONAL LABORATORIES
Division of Richardson-Merrell Inc.
Cincinnati, Ohio 45216
6-8888 (area code)

Nothing motivates like early weight loss

Help
motivate
with

Tenuate® (diethylpropion hydrochloride N.F.)

Merrell

Wednesday, February 5, 1975

MEDICAL TRIBUNE

One Man... and Medicine

ARTHUR M. SACKLER, M.D.,
International Publisher, Medical Tribune



Wanted: Reasons Why

Brinkmanship was no monopoly of John Foster Dulles.

Hundreds of millions throughout the world teeter on the brink of starvation; yet social leaders focus on dangers of the year 2,000 instead of getting food to those starving now. Why? Scores of millions hover on the brink of health and die from the ravages of preventable and treatable diseases—yet confidence in doctors and their drugs are constantly undermined. Why? Huge gaps exist in

our knowledge—gaps that must be closed through new knowledge if, surviving the threat of war and hunger, we are to progress to new levels of health; yet major sectors of biomedical research are under attack now. Why?

One reads with horrified fascination of the well-intentioned but potentially disastrous efforts of those who would deprive prisoners of their social right to volunteer as subjects for research. Why? Why don't those opposing research on mental patients explain how in heaven's name we are going to help these people with better therapies than those which are presently available?

What are the ulterior motives? Are they protection of personal credits and dogmas and the imposition of these on others? Are they the need to create issues by those seeking political and social change? Do they reflect new forms of personal gain—rewards in the coinage of publicity and press prominence?

Misleading Distortion

One can accept the right of individuals to defend their own beliefs but not necessarily to impose them on others. One can defend the right of individuals who seek social change. But one cannot accept the misleading distortion of issues for disguised political or personal motives, for publicity or prestige.

It is tragic that the opportunity to do biomedical research is being undermined, the freedom with which to disseminate its findings restricted, and the time necessary for its application in the biomedical area being constantly lengthened. The crusaders of our day, whether for religious or consumer advocacy, have learned the power of publicity pressure in the political arena and on the bureaucracies of government.

The threats to the biomedical sciences are like those of a multi-headed hydra—one sooner is one chopped off than one or two more appear. The Department of Agriculture has restricted the import of animals, including higher primates, with senseless disregard of the implications for therapeutic and basic research. Why? Government regulatory actions constantly proliferate more and more restrictions without regard to compensatory gains. Why? Simplistic slogans and similes are proposed without basis in experience or study but apparently primarily on "the courage of their confusions."

Why the continuing escalation of attacks on medicine and men of medicine? Why the ever-increasing threats to research in the biomedical sciences? Why have we progressed so little in shortening the interval between discovery and application in this, the day of instantaneous communication and mass education? Why do we revert again and again to the earlier periods of anti-science when anatomy depended upon stolen bodies and when the efforts of scientists were challenged by the dogmas of established beliefs?

The witch hunters of an earlier day burned the bodies of their victims

GE Technique Spots Subtle Heart Defects



Heart defects undetected by routine electrocardiograms may now be identified by a technique being developed by G.E. The technique combines a mini-computer with a supersensitive electronic "ear" that provides a much broader and more accurate range of heart sounds, which are computer-analyzed on the spot for interpretation and diagnosis.

For UN Staffers in Geneva, Blues Often Mar Blue Skies

Medical Tribune World Service

GENEVA—In the eyes of many Swiss, members of the international staff of the United Nations Organization lead an enviable existence, with high salaries, no income tax, and certain diplomatic privileges, including cheap liquor and gasoline and virtual immunity to parking tickets.

But, in fact, emotional problems are common among these international careerists. The single woman, for instance, may have to cope with loneliness in a huge faceless organization, along with the difficulties of adapting to an alien culture.

When the menopause approaches, women in this situation may suffer from depression to an unusual degree. MEDICAL TRIBUNE was told in an interview here by Dr. Jean-Felix Dulac, head of the U.N. medical service. He cited cases of alcoholism, and attempted suicide.

Stresses May Be Severe

Dr. Dulac pointed out that the stresses incident to taking a job with a U.N. organization may be severe, and even top executives may need as much as two years before settling down to effective work.

The stresses are not confined to the U.N. organizations in Switzerland. Dr. Dulac noted that they are also common in staff in New York and Paris.

One difficulty in treatment of sufferers is language. In Geneva, for example, many psychiatrists and other psychotherapists are fluent in English (and often German and Italian) as

High BP Found in 42 %

Medical Tribune Report

NEW YORK—A recent survey of 1,545 passers-by in the lobby of the Empire State Building has shown that "42 per cent of the New York population is walking around with high blood pressure," the Preventive Medicine Institute—Strang Clinic has reported.

Almost half (45 per cent) of men between the ages of 40 and 64 had blood-pressure readings above normal, and nearly 40 per cent of women in the same age-group had readings that "would require medical attention," the survey showed. In the under-40 group, women fared much better than men.

The Pseudo-ulcer



Ulcer-like symptoms: no G.I. pathology

The patient is convinced it's an ulcer. However, symptoms are not quite typical, and x-ray findings are negative. These findings and the results of additional diagnostic procedures exclude an organic basis for the patient's complaints. A diagnosis of "upper functional gastrointestinal disorder" is made, which is supported by the fact that episodes of painful symptoms coincide with episodes of excessive anxiety, as indicated by the history.

It may be useful to explain to the patient the mechanism by which emotions upset normal G.I. functioning, resulting in hypersecretion and hypermotility and thus causing such symptoms as nausea and epigastric pain. In upper functional gastrointestinal disorders, counseling by the primary physician can often help the patient to understand how excessive anxiety may cause flare-ups of G.I. symptoms.

A disproportionate number of patients seen by the general practitioner suffer from functional disorders, as do more than half of those seen by the gastroenterologist.* Where milder cases may respond to counsel-

ing alone, if symptoms are severe and disabling to any degree, a suitable regimen may include medication to reduce the symptoms and the excessive anxiety that often provokes these distressing symptoms. In these cases, Librax as an adjunct can greatly contribute to the course of therapy. Its dual action can offer relief of both painful symptoms and excessive anxiety, because each capsule contains 5 mg chlordiazepoxide HCl and 2.5 mg clidinium Br. The antianxiety action of Librium® (chlordiazepoxide HCl) makes Librax exceptional among drugs for certain gastrointestinal disorders associated with excessive anxiety; the clidinium bromide (Quarzan®) component furnishes dependable antisercretory-antispasmodic action. Dosage is flexible; it may be adjusted according to your patient's requirements within the range of 1 or 2 capsules three or four times daily, up to 8 capsules daily in divided doses.

*Rome HP, Brannick TL: Orientation and mechanism of functional disorders: clinicophysiological correlation, chap. 153, in *Gastroenterology*, edited by Bockus HL, Philadelphia, WB Saunders Company, 1965, p. 1110.

An adjunct
in anxiety-related upper
functional G.I. disorders

Librax®

Each capsule contains 5 mg chlordiazepoxide HCl and 2.5 mg clidinium Br.

Before prescribing, please consult complete product information, a summary of which follows:

Indications: Symptomatic relief of hypersecretion, hypermotility and anxiety and tension states associated with organic or functional gastrointestinal disorders; and as adjunctive therapy in the management of peptic ulcer, gastritis, duodenitis, irritable bowel syndrome, spastic colitis, and mild ulcerative colitis.

Contraindications: Patients with glaucoma; atrial fibrillation; benign bladder neck obstruction; known hypersensitivity to chlordiazepoxide hydrochloride and/or clidinium bromide.

Warnings: Caution patients about possible combined effects with alcohol and other CNS depressants. As with all CNS-acting drugs, caution patients against hazardous occupations requiring complete mental alertness (e.g., operating machinery, driving). Though physical and psychological dependence have rarely been reported on recommended doses, use caution in administering Librax (chlordiazepoxide hydrochloride) to known addiction-prone individuals or those who might increase dosage; withdrawal symptoms (including convulsions), following discontinuation of the drug and similar to those seen with barbiturates, have been reported. Use of any drug in

pregnancy, lactation, or in women of childbearing age requires that its potential benefits be weighed against its possible hazards. As with all anticholinergic drugs, an inhibiting effect on lactation may occur.

Precautions: In elderly and debilitated, limit dosage to smallest effective amount to prevent development of ataxia, oversedation or confusion (not more than two capsules per day initially; increase gradually as needed and tolerated). Though generally not recommended, if combination therapy with other psychotropics seems indicated, carefully consider individual pharmacologic effects, particularly in use of potentially drug such as MAO inhibitors and phenothiazines. Observe usual precautions in presence of impaired renal or hepatic function. Paradoxical reactions (e.g., excitement, stimulation and acute rage) have been reported in psychiatric patients. Employ usual precautions in treatment of anxiety states with evidence of superimposed depression; suicidal tendencies may be present and protective measures necessary. Variable effects on blood coagulation have been reported very rarely in patients receiving the drug and oral anticoagulants; causal relationship has not been established clinically.

Adverse Reactions: No side effects or manifestations not seen with either compound alone have been reported with Librax. When chlordiazepoxide hydrochloride is used alone, drowsiness,



Roche Laboratories
Division of Hoffmann-La Roche Inc.
Nutley, New Jersey 07110

The Only Independent Weekly Medical Newspaper in the U.S.

Medical Tribune

and Medical News
Published by Medical Tribune, Inc.

Some News Items . . .

ITEM #1—Blue Cross-Blue Shield of Greater New York is asking for a 27.8 per cent average rate increase for hospital charges to go into effect March 1, 1975. The last increase totalled 7.4 per cent on April 1, 1973. Of the present increase, 22.2 per cent will be for costs of current benefits and 5.6 per cent for proposed new benefits.

ITEM #2—The increases are attributed to "rising prices for food, fuel and other

supplies, and higher collectively bargained hospital wages, salaries and fringe benefits."

ITEM #3—New York City's Health and Hospitals Corporation is seeking to raise its present per diem from Blue Cross of \$117 a day for in-patient hospitalization to as much as \$200 a day, reportedly comparable to reimbursement rates in private and voluntary institutions.

... Relevant Queries . . .

DO PRESENTLY proposed government health insurance plans project this rate of inflation and per diem hospitalization at almost \$200? How will the government meet "rising prices for food, fuel and other supplies, and higher collectively bargained hospital wages, salaries and fringe benefits"? Cutting costs of drugs and doctors' fees will not suffice. As inflation escalates in cost, will services be cut and availability and duration of hospitalization restricted for beneficiaries of federal programs? Are government projections for

administrative costs comparable to those of the Blue Cross and Blue Shield plans? What are these projections and when were they made?

How long will it take our governmental agencies to realize that no national health insurance program will be viable without massive expansion of health manpower and preventive medicine; without more effective action in respect to addicting cigarettes and alcohol, and without the development of new medicines to reduce both the need for and the duration of hospitalization?

... And Some Major Questions

GOOD preventive medicine, more new medicines and earlier diagnosis of treatable disorders are realistic national needs; not rhetorical posturing and phony bureaucratic "cost effectiveness" proposals. America has seen the type of bureaucratic regulation which has virtually destroyed the American railroad system and has crippled our postal service; such bureaucratization can

also bankrupt or cripple our presently functioning, albeit not perfect, health care distribution system.

Why don't our "double-blind" health officials who require well-controlled experiments for individual drugs and devices test their proposals? Why are there no prototype pilot projects to check the validity of their proposed changes in our health care system? A.M.S.

Where Are All The Unmarried Men?

THE ABOVE question is taken from an article in a recent issue of the *Statistical Bulletin* and refers solely to Americans. On the basis of the 1970 census of the population, the number of unmarried men aged 18-29 per 100 unmarried women aged 16-24 is 110 in the Pacific states; it is 104 in the South Atlantic states. In the remaining seven geographic subdivisions of the U.S., unmarried men are outnumbered by unmarried women with the greatest discrepancy occurring in the East North Central states where there are 88 such men for every 100 such women.

unmarried men per 100 unmarried women, such as 213 in Alaska, 146 in Hawaii, 120 in Nevada, 120 in Rhode Island, 115 in Virginia, 112 in South Carolina and 111 in California. Unmarried women exceed unmarried men by more than 15 per cent in Minnesota, Pennsylvania, Ohio, Iowa, West Va. and Utah.

How does one explain these geographical concentrations of single men? Alaska is our last frontier but that certainly is not true of Hawaii, Rhode Island or Virginia.

Perhaps now that the word is out the ratios will be readjusted.

Emergency Medical Service

CLINICAL QUOTE: "You can't predict when you go out on an emergency which call will need advanced life support. The older patient who falls and breaks a hip may have done so because of arrhythmia. A heart problem may cause an automobile accident, and then arrhythmia complications may lead to cardiac arrest en route to the

hospital." (Dr. Costas T. Lambrew, Chairman, Department of Medicine at Nassau County Medical Center, Long Island, N.Y., after analyzing the EKG's of 9,000 patients—1,728 with chest pain, 4334 with illnesses other than chest pain, 2744 trauma victims and 194 unclassified patients. See page 1.)



"Please make the check payable to Dr. Jekyll and/or Mr. Hyde."

©1975 Medical Tribune

LETTERS TO TRIBUNE

His Own Spokesman

In a recent editorial (MT, Dec. 4, 1974) you asserted that the A.M.A. has long been the "official spokesman" of medicine, advocating the views of the majority of its members. That is much like saying a labor union or a government represents the views of the majority. This is not true. In fact, it is fraud to perpetuate the myth of any person or group "representing" a given individual. Only I can represent my views. I might give another person the authority to represent my views on a certain matter, but there is no way he can "represent" me in a broad sense. Therein lies the futility and fallacy of democracy, voting, government or coercion in any form.

It is time we recognize the individual. He is the possessor of inalienable rights to his life, liberty and property. He and he alone is sovereign. This is one of the most potent facts of life. If only enough men will act according to their nature qua man and abhor the tribe and all forms of collectivism, altruism and sacrifice, it could be a better world.

All any man has to do is eliminate evil. He is the possessor of inalienable rights to his life, liberty and property. He and he alone is sovereign. This is one of the most potent facts of life. If only enough men will act according to their nature qua man and abhor the tribe and all forms of collectivism, altruism and sacrifice, it could be a better world.

ROBERT S. BORDEN, M.D.
Groton, Mass.

personnel and sophisticated laboratory facilities to perform tests the private physician requires for diagnostic confirmation. The headquarters of the National Genetics Foundation acts as a clearinghouse by directing physicians and/or the lay public to the appropriate medical center with the most comprehensive facilities for the particular problem involved.

In the past four years physicians associated with hospitals throughout the country have been utilizing this important service which is often vital to those physicians involved in the practice of pediatrics, obstetrics, or family medicine. Services are secured by contacting the National Genetics Foundation directly at the above address, or by telephone.

GEORGE W. MELCHER, JR., M.D.
New York, N.Y.

Reviewing by What Peers

I could not agree more with the letter by Dr. James K. Theisen (MT, Dec. 18, 1974) concerning the fact that PSRO has not been accepted—nor should it be. Congress itself has two bills pending concerning the repeal of this law and each of us should write our congressmen requesting action on these bills—HR 12256 (Mr. Rarick, Mr. Parry, Mr. Lott, Mr. Flynn—Jan. 23, 1974) and HR 15266 (Mr. Brody—June 6, 1974). Nothing can be changed if we don't move to change it ourselves.

H. TAYLOR YATES, JR., M.D.
Alexandria, Va.

Genetic Counseling

I read with interest Dr. Kurt Hirschhorn's "In Consultation" article, "What is New and Important in Genetic Counseling?" (MT, Dec. 18, 1973).

In regard to where to turn for genetic counseling help, I would like to call your attention to the National Genetics Foundation, (250 West 57th Street, New York, N.Y. 10019, (212-265-3166) which offers a unique service to the physician by providing assistance with any genetic or genetically related problem. The National Genetics Foundation operates a network of genetic centers involving 47 medical teaching institutions throughout the United States and Canada. Many of its participating centers have the trained

Dr. Coolidge's Tube

I was pleased to see you publish a picture, and to know that Dr. William D. Coolidge is still alive. As you well know, the invention by Dr. Coolidge of the hot cathode Roentgen tube with an electrically heated cathode permitted close and careful regulation of the quality and quantity of X rays emitted by the tube. Prior to his invention, the old gas tube was very unreliable and unpredictable, and could not be carefully calibrated.

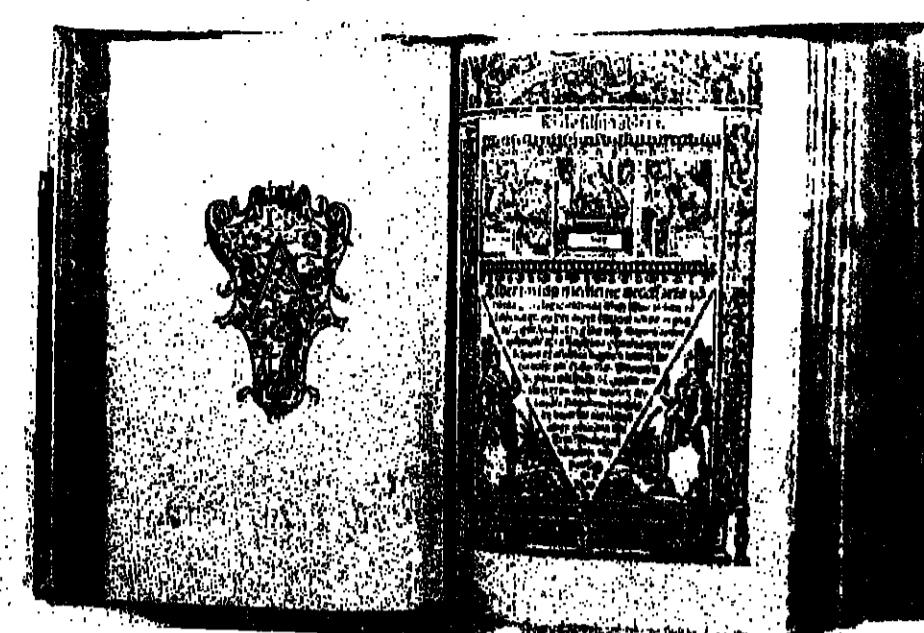
This remarkable man made Roentgen's invention practical.

Many thanks for your fine publication.

D. F. CAPPARATTI, M.D.
Fallon, Nev.

Penn Hospital Library Gets Refurbishing

PENNSYLVANIA Hospital in Philadelphia has the country's oldest medical library, formally founded in 1726. The library contains what is believed to be the most extensive medicohistorical collection owned by a hospital and one that is distinctive in its continuity as a working medical library for the century 1752-1852. The hospital is using a grant from the Department of Health, Education, and Welfare, the Public Health Service, and the National Library of Medicine to reorganize the archives. Shown here are some items from the collection.



Wednesday, February 4, 1975

Wednesday, February 5, 1975

Cover from *Articella Medicatio*, by Cypriano Heyll, written in 1534. The book discusses the works of Claudius Galen. At right is the cover page from a medical treatise written in 1523 and bound within the book.



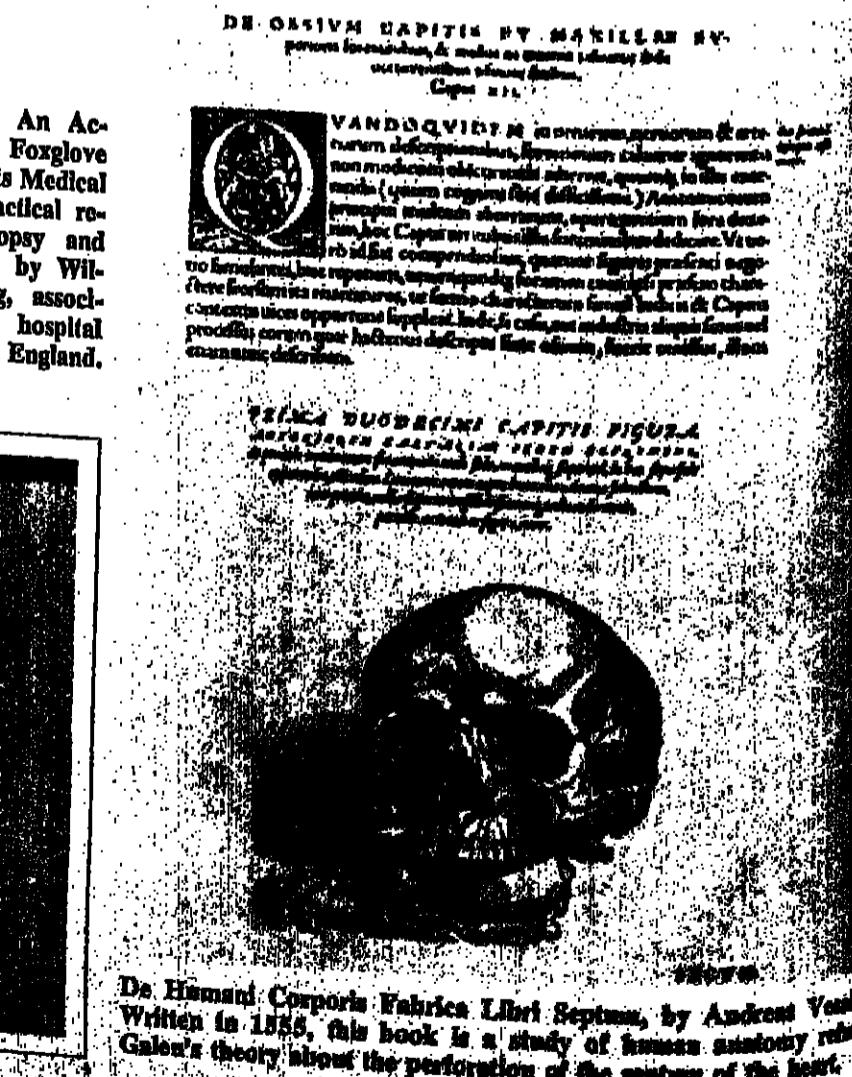
1970: coronary care unit.

Human Spirit's Triumph Depicted In Medical Center Photo Exhibit

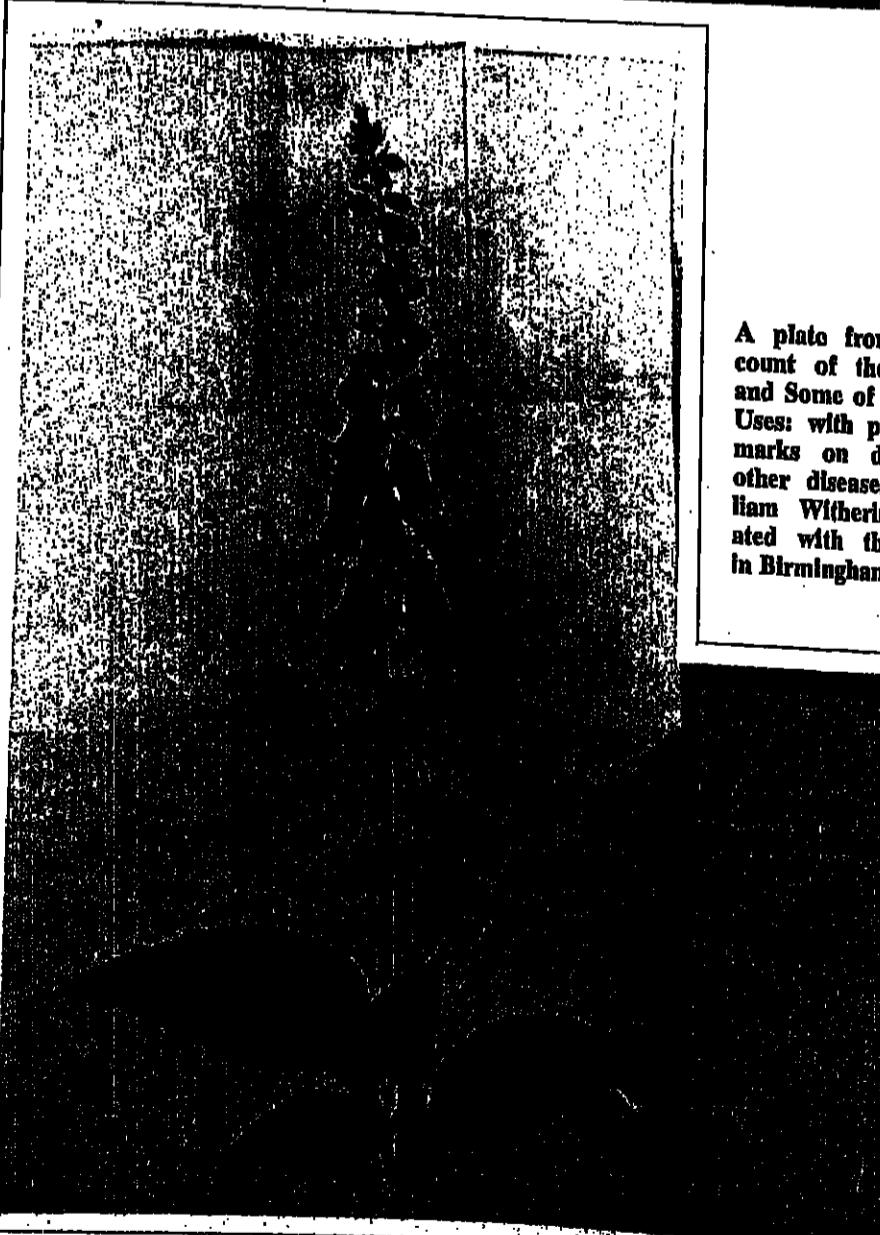
THE MEDICAL CENTER PERCEIVED," a major exhibition of photographs taken at the Albany Medical Center by photographer Dan Budnik during the years 1959-74, was recently on display at the Art Gallery of the State University of New York at Albany. Sponsored by the center in observance of the 125th anniversary of the founding of Albany Medical Center Hospital, the 134 photographs selected from among 19,000 that Mr. Budnik took over the 15-period reflect the theme of the triumph of the human spirit in the face of pain and adversity. Shown here are a few of the photos from the exhibit.



Library personnel and students are working, left, to dust, inventory, and code as well as catalogue the library's collection. The library contains 15,000 volumes as well as paintings and documents.



De Oestivis Capitis by Macmillan 1810. William Withering, an English physician, made an extensive study of foxglove and its medical uses. His book, 'An Account of the Foxglove and Some of its Medical Uses: with practical remarks on dropsy and other diseases,' was published in 1785. Withering, associated with the hospital in Birmingham, England.



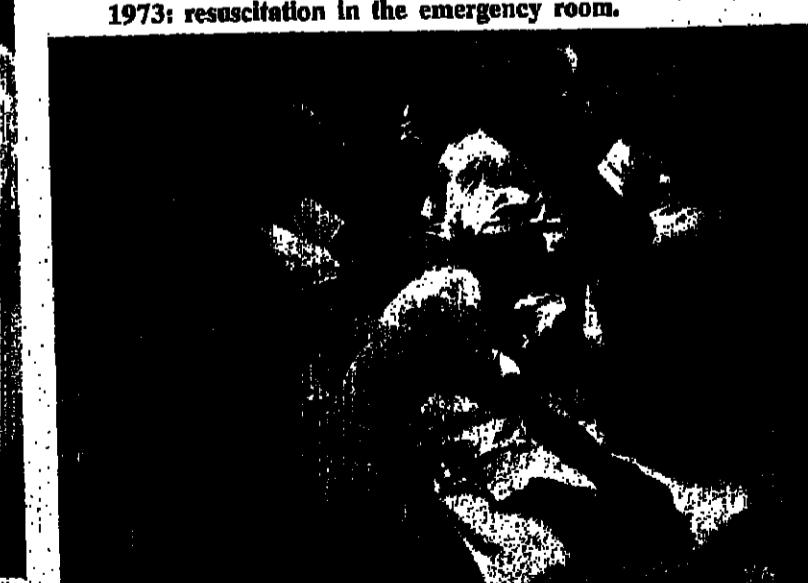
De Humani Corporis Fabrica Libri Septem, by Andreas Vesalius. Written in 1543, this book is a study of human anatomy. Vesalius' theory about the perforation of the septum of the heart.



1962: heart surgeon.



1974: burn unit.



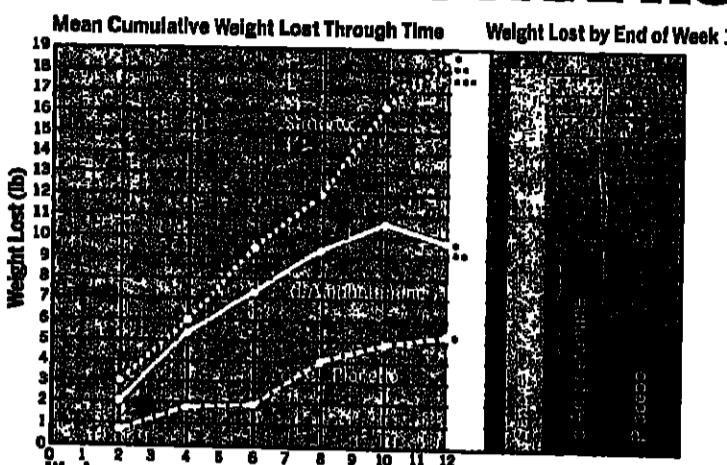
1973: resuscitation in the emergency room.

1965: freshmen medical students in neuroanatomy laboratory.

SANOREX® IN OBESITY (MAZINDOL)®

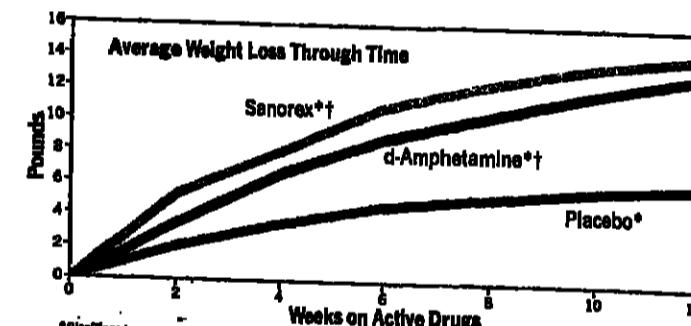
TABLETS, 1 mg and 2 mg.

AS EFFECTIVE AS d-AMPHETAMINE



In a double-blind study² of 40 obese patients (all of whom completed the study), Sanorex (1 mg t.i.d.) was more effective than either placebo or d-amphetamine (5 mg t.i.d.) in helping patients lose weight.

The 14 patients on Sanorex experienced a substantially greater mean weight loss—1½ to 2 lb/wk, as compared with 1 to 1½ lb/wk for the 14 d-amphetamine patients—throughout the 12-week phase of active medication. After the sixth week, the superiority of Sanorex became increasingly evident. And as treatment progressed, so did weight loss in patients on Sanorex—whereas after the tenth week, patients on d-amphetamine began to regain some weight.



In a double-blind study³ of 93 obese patients (all of whom completed the study), 30 patients received Sanorex (1 mg t.i.d.), 31 received placebo, and 32 received d-amphetamine (5 mg t.i.d.).

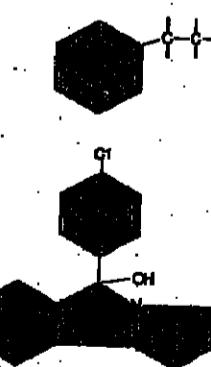
During the 12-week phase of active medication, patients on Sanorex lost an average of 14.1 lb, compared with 13.1 lb for d-amphetamine patients and 5.6 lb for placebo patients. Throughout the active medication phase, 63% of patients on Sanorex lost more than 1 lb/wk, compared with 38% of the d-amphetamine group and 29% of the placebo group.

BUT WITH CERTAIN DIFFERENCES

Although the pharmacologic activity of Sanorex and that of amphetamines are similar in many ways (including central-nervous system stimulation in humans and animals, as well as production

of stereotyped behavior in animals), animal experiments suggest that there are differences.⁴ Sanorex also differs in basic chemical structure from amphetamines and all other prescription anorectants.

Different Chemical Structure



An important chemical similarity between amphetamines and all other prescription anorectants except Sanorex is the basic phenethylamine structure to which their differentiating chemical radicals are attached.

An important chemical difference between Sanorex and all other prescription anorectants is that Sanorex is an isoindole; it does not contain a phenethylamine structure.

Simplicity and Flexibility of Dosage

Simple one-a-day dosage is facilitated by 2-mg tablets (taken 1 hour before lunch).

New flexibility (for the patient in whom 1 mg t.i.d. is preferred) is now facilitated by new 1-mg tablets (taken 1 hour before meals).

For Brief Summary, please see facing page.

SANOREX® (MAZINDOL)®

References
 1. Kornheber A: Problems and current concepts in the treatment of obesity. Scientific Exhibit presented at the New York State Academy of Family Physicians 25th Annual Scientific Convention, McAfee, NJ, May 8-10, 1973.
 2. DeFelice EA, Cheykin M, Cohen A: Double-blind clinical evaluation of mazindol, dexamphetamine, and placebo in treatment of exogenous obesity. *Am J Clin Ther* 15:358-365, July 1973.
 3. Vernace B: Practical considerations for managing obese patients: Initial interview and affective treatment in the office. Scientific Exhibit presented at the American Medical Association, 27th Clinical Convention, Anaheim, Calif, Dec 1-4, 1973.

Indication: In exogenous obesity, as a short-term (a few weeks) adjunct in a weight reduction regimen based on caloric restriction. The limited usefulness of agents of this class should be measured against possible risk factors.

Contraindications: Glaucoma; hypertension or idiosyncrasy to the drug; agitated states; history of drug abuse; during, or within 14 days following, administration of monoamine oxidase inhibitors (hypertensive crisis may result).

Warnings: Tolerance to many anorectants may develop within a few weeks; if this occurs, do not exceed recommended dose, but discontinue drug. May impair ability to engage in potentially hazardous activities, such as operating machinery or driving a motor vehicle, and patient should be cautioned accordingly.

Drug Interactions: May decrease the hypotensive effect of guanethidine; patients should be monitored accordingly. May markedly potentiate pressor effect of exogenous catecholamines; if a patient recently taking mazindol must be given pressor amine agents (e.g., levaterenol or isoproterenol) for shock (e.g., from a myocardial infarction), extreme care should be taken in monitoring blood pressure at frequent intervals and initiating pressor therapy with a low initial dose and careful titration.

Drug Dependence: Mazindol shares important pharmacologic properties with amphetamines and related stimulant drugs that have been extensively abused and can produce tolerance and severe psychological dependence. Manifestations of chronic overdose or withdrawal with mazindol have not been determined in humans. Abstinence effects have been observed in dogs after abrupt cessation for prolonged periods. There was some self-administration of the drug in monkeys. EEG studies and "liking" scores in human subjects yielded equivocal results. While the abuse potential of mazindol has not been further defined, potential of dependence should be kept in mind when evaluating the desirability of including the drug in a weight-reduction program.

Usage in Pregnancy: In rats and rabbits an increase in neonatal mortality and a possible increased incidence of rib anomalies in rats were observed at relatively high doses. Although these studies have not indicated important adverse effects, the use of mazindol in pregnancy or in women who may become pregnant requires that potential benefit be weighed against possible hazard to mother and infant.

Usage in Children: Not recommended for use in children under 12 years of age.

Precautions: Insulin requirements in diabetes mellitus may be altered. Smallest amount of mazindol feasible should be prescribed or dispensed at one time to minimize possibility of overdose. Use cautiously in hypertension, with monitoring of blood pressure; not recommended in severe hypertension or in symptomatic cardiovascular disease including arrhythmias.

Adverse Reactions: Most commonly, dry mouth, tachycardia, constipation, nervousness, and insomnia.

Cardiovascular: Palpitation, tachycardia.

Central Nervous System: Overstimulation, restlessness, dizziness, insomnia, dysphoria, tremor, headache, depression, drowsiness, weakness.

Gastrointestinal: Dryness of mouth, unpleasant taste, diarrhea, constipation, nausea, other gastrointestinal disturbances.

Skin: Rash, excessive sweating, clamminess.

Endocrine: Impotence, changes in libido have rarely been observed.

Eye: Long-term treatment with high doses in dogs resulted in some corneal opacities, reversible on cessation of medication; no such effect has been observed in humans.

Dosage and Administration: 1 mg three times daily, one hour before meals, or 2 mg per day, taken one hour before lunch in a single dose.

How Supplied: Tablets, 1 mg and 2 mg, in packages of 100.

Before prescribing or administering, see package circular for Prescribing Information.

24-2610 SANOFI

SAUNDERS PHARMACEUTICALS, EAST HANOVER, N.J. 07933

New US Role Offers Hope on Traffic Deaths

Continued from page 1
period in 50-50 matching grants for planning, implementation and research.

For the fiscal year ending June 30, 1974, the fund underwrote 88 projects for a total federal share of \$27 million. For approval, grant applications must outline plans for EMS components covering both pre-hospital and post-hospital phases. Without the systems approach, EMS planning tended to be fragmented, according to Dr. Boyd. "Most places just bought ambulances through the Department of Transportation without putting in the total program," he points out. "But, now, with our money, they can take on the total comprehensive package, including ambulances bought by DOT."

Turning Point Seen

Although it's too soon for the projects to have made any headway, Dr. Boyd is confident the new federal role is a significant turning point. "I think this is what the country has been waiting for," he says. "It's the first time there's been a lead agency and a planned program built on patient care necessity. We've finally got someone up here where the buck stops. We've long had the expertise, the learning experience. Now we can transpose that."

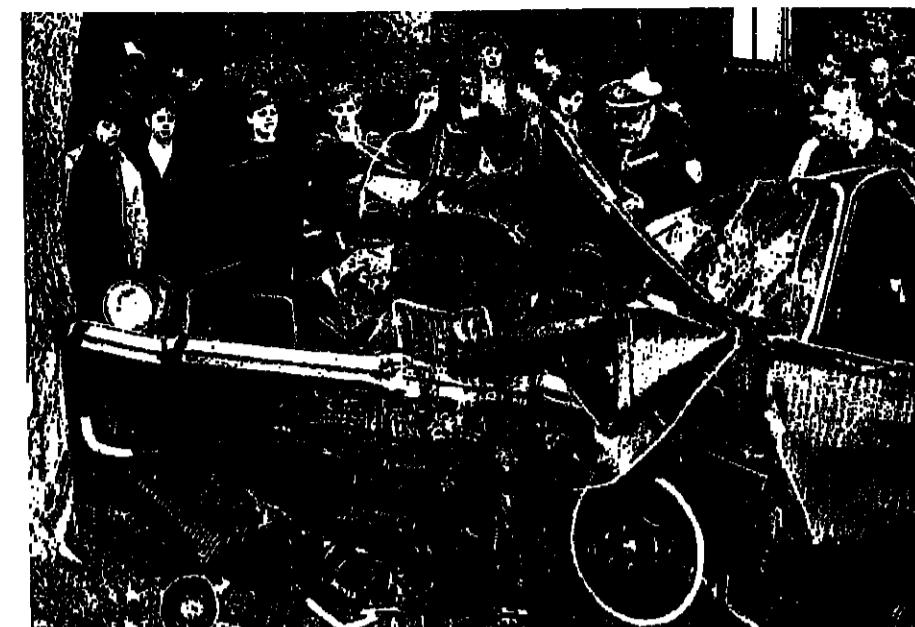
Where to start is the question facing many communities. Some observers, pointing to the fact that at least half of all heart attack and accident victims die before they reach the hospital, argue that communications and transportation are the logical priorities. "If something isn't done in the first four minutes," notes Jerry Montgomery, Director of Emergency Services at Evanston Hospital in Evanston, Illinois. "It may be the fire department, the police, or the ambulance service, as long as the number is highly visible to the public."

Ambulances. According to an HEW survey in mid-1971, 44 per cent of 25,000 ambulances in 37 states were operated by funeral homes. Dr. Boyd reckons that the figure hasn't changed much and may actually be as high as 80 per cent in rural areas. In tiny McCormick, South Carolina, with a population of 2,000, only a flashing red light on top of the vehicle distinguishes the ambulance from a hearse. "If they switch it off halfway to the hospital," reports a local physician, "you know what happened."

Pre-Hospital Phase Lags

In Illinois, which has developed what is probably the most sophisticated state-wide EMS system in the country, the focus was on the hospitals and as a result, the service still lags behind in the upgrading of the pre-hospital phase. In a much-publicized case that occurred in January, a pregnant woman died of a massive blood clot at a hospital just three blocks from her home when the local fire chief judged the situation a non-emergency and transferred the request for an ambulance to a fire department further away.

"We have to get them into the system sooner," emphasizes Blair L. Sadler, assistant vice president of the Robert Wood Johnson Foundation of Princeton, N.J., which in May announced 44 demonstration grants totaling \$1.5 million for projects with attention directed toward access, training of personnel and a central dispatch facility.



That deaths from traffic accidents dropped in 1974 is attributed to the 55-mile speed limit rather than emergency medical services.

the exorbitant expense of a telemetry system effectively," comments Dr. Clarence Hart, an orthopedic surgeon and president of the Illinois division of The American Trauma Society. "You need to have a large population concentrated in a small area. I don't think the effort will hold up in the rural areas."

Arrhythmias Common

According to a recent study, it may be propitious for any community to consider the added expense of telemetry, and not only for cardiac victims. When Dr. Costas T. Lambrew, Chairman of the Department of Medicine at Long Island's Nassau County Medical Center, analyzed the EKG's of 9,000 patients—1,728 with chest pain, 4,334 with illnesses other than chest pain, 2,744 trauma victims and 194 unclassified—he made some surprising discoveries (*Heart & Lung*, Sept, Oct 1974). Significant arrhythmias were documented in 8.7% of patients with complaints other than chest pains. In some instances, arrhythmia was responsible for the symptoms and knowledge edge for its presence was vital in the immediate care of the patient. In still others, documentation of an arrhythmia, even of one not an immediate threat to life, was found to be important in proper medical care of the patient.

"You can't predict when you go out on an emergency which call will need advanced life support," Dr. Lambrew concludes. "The older patient who falls at home and breaks a hip may have done so because of arrhythmia. A heart problem may cause an automobile accident, and then arrhythmia complications may lead to cardiac arrest en route to the hospital."

38 Frequencies Allocated

With the allocation by the Federal Communications Commission of 38 frequencies for emergency medical use under the EMS Systems Act, ambulance radio communication should improve, although many ambulances still lack radios.

Expensive

Expensive telemetry monitoring systems which are being installed in many new EMS operations, have made some experts uneasy, especially about their use in rural areas. "In order to support

The status of paramedic personnel—requiring an additional 120 hours of

Continued on page 29

Only one antihypertensive provides the three preferred modes of action.

Ser-Ap-Es®

reserpine 0.1 mg
hydralazine hydrochloride 25 mg
hydrochlorothiazide 15 mg

INDICATIONS
Based on a review of this drug by the National Academy of Sciences-National Research Council, and information, FDA has classified the indications as follows:

Effective: Hypertension. (See box warning.)

WARNING
This fixed combination drug is not indicated for the therapy of hypertension. Hypertension requires individualized therapy to the individual patient. If the fixed combination represents the dosage so determined, its use may be more convenient in patient management. The control of hypertension is not static, but must be reevaluated as conditions in each patient warrant.

CONTRAINDICATIONS
Reserpine: Known hypersensitivity; mental depression (especially with suicidal tendencies); active tuberculosis; ulcerative colitis; electroconvulsive therapy.

Hydralazine: Hypersensitivity; coronary artery disease; mitral valvular rheumatic heart disease. Hydrochlorothiazide: Anuria; hypersensitivity to thiazides; sulfonamide-derived drugs. The routine use of diuretics in healthy pregnant women with or without mild edema is contraindicated and possibly hazardous.

WARNING
Reserpine: Use with extreme caution. In patients with a history of mental depression. Discontinue at first sign of depression, early morning insomnia, loss of appetite, impotence, or mild depression. Drug-induced depression may persist for several months after drug withdrawal and may be severe enough to lead to suicide. MAO inhibitors should be avoided or used with extreme caution.

Hydralazine: Acute administration of doses over 400 mg daily may cause an syndrome simulating acute systemic vasculitis (erythema). This may also occur at lower doses. Long-term treatment with steroids may be necessary to control this syndrome which may be manifested years later. CSECA, liver function, and antinuclear antibody titer determinations are indicated before and periodically during prolonged therapy with hydralazine if the patient develops any unexpected signs or symptoms. Use MAO inhibitors with caution.

Hydrochlorothiazide: Use with caution in severe renal disease. In patients with renal disease, the drug precipitates azotemia. Cumulative effects of the drug may develop in patients with impaired renal function.

Thiazides should be used with caution in patients with impaired hepatic function or progressive liver disease, since minor alterations of fluid and electrolyte imbalance may precipitate hepatic coma.

Thiazides may be additive or potentiate the action of other antihypertensive drugs. Potentiation of antihypertensive drugs. Potentiation of adrenergic blocking drugs.

Sensitivity reactions are more likely to occur in patients with a history of allergy or bronchial asthma.

The possibility of exacerbation or activation of systemic lupus erythematosus has been reported.

Usage in Pregnancy

Reserpine: The safety of reserpine for use during pregnancy or lactation has not been established. Patients on reserpine should be weaned off the drug as soon as possible. It is essential to the welfare of the patient, if increased respiratory tract secretions, nasal congestion, and breastfed infants are exposed to reserpine mothers since reserpine crosses the placental barrier and appears in maternal breast milk.

Hydralazine: The drug should be used only when, in the judgment of the physician, it is deemed essential to the welfare of the patient.

Hydrochlorothiazide: Usage of thiazides in women of childbearing age requires that the potential benefits of the drug be weighed against its possible hazards to the fetus. These hazards include fetal or neonatal syndrome, thrombocytopenia, and possibly other adverse reactions which have occurred in the adult.

Nursing Mothers
The drug crosses the placental barrier and appears in cord blood and breast milk.

PRECAUTIONS

Reserpine: Use cautiously in patients with hypertension, glaucoma, ulcerative colitis, or gallstones (follicular colitis may be precipitated).

Exercise caution when treating hypertension with renal insufficiency. Use cautiously with digitalis and quinidine.

Intravenous hypotension has occurred in patients receiving intravenous preparations, but withdrawal of reserpine does not assure that circulatory instability will not occur in such patients.

Hydralazine: Use cautiously in suspected coronary artery disease, peripheral vascular disease, cerebral vascular accidents, and nephritis, renal damage. Postural hypotension may occur, and the pressor response to epinephrine may be reduced.

In treating hypertension, current clinical practice stresses the importance of achieving control of three basic homeostatic mechanisms: fluid volume, sympathetic activity, and arteriolar tone.¹

Initial treatment most frequently employs one of the thiazides.²⁻⁷

But if blood pressure resists fluid volume control with thiazides, second agent with a different mode of action, such as a sympathetic inhibitor (reserpine), may be gradually added.⁸⁻¹¹

Many hypertensives, however, may resist control even with a two-drug regimen.

In such cases, the crucial "third step" in combined therapy is frequently control of arteriolar tone with hydralazine.²⁻⁴

Ser-Ap-Es combines all three steps in a single tablet—all the medication many hypertensives will need.

And when the dosage of each component corresponds to the dosages pre-established by individualized titration, Ser-Ap-Es may prove more convenient and more economical.

Doses of each component in Ser-Ap-Es are lower than when used alone.

Note: Use Ser-Ap-Es cautiously in patients with advanced renal damage or cerebrovascular accident. Discontinue at first sign of mental depression.

Ser-Ap-Es is the only antihypertensive agent that provides the three basic drugs used in two published VA cooperative studies.^{8,9}

References: 1. Freis ED: Hypertension: a curable disease. *Clin Pharmacol Ther* 13:627-632, 1972. 2. Even a little is too much. *Emergency Med* 8:144-185, 1973. 3. Bender AD, Familiar RD: Combination drugs in the treatment of hypertension. In: *Management of Essential Hypertension*. Falon JL (ed): *Practical Clinical Hypertension*. In: *Harrison's Principles of Internal Medicine*, ed 18. New York, Appleton-Century-Crofts, 1972, pp 33-40. 4. Gifford RW Jr: Drugs for arterial hypertension. In: *Drugs of Choice*, ed 1. St. Louis, The CV Mosby Co, 1972, pp 393-395. 5. Sellers AM, Itskovitz H: Inderal: a new antihypertensive agent. *Arch Intern Med* 131:1371, vol 11, pp 934-943. 6. Conn HL Jr, Horwitz O (eds): *Cardiac and Vascular Disease*. Philadelphia, Lippincott, 1971. 7. Effects of treatment on blood pressure in hypertension. Results of the Veterans Administration Cooperative Study Group on Antihypertensive Agents. *JAMA* 202:1028-1034, 1984. 8. Effects of treatment on average blood pressure. 14 mm Hg. Veterans Administration Cooperative Study Group on Antihypertensive Agents. *JAMA* 213:1143-1152, 1970.

Peripheral neuritis, evidenced by paresthesias, numbness, and pain, has been observed.

Published evidence suggests a hypothyroid effect and addition of thyroxine to the regimen if symptoms develop.

Bradycardia, constipation, as with reduction in hemoglobin and hematocrit, and peripheral edema, as with hydralazine, and purpura, have been reported. If such abnormalities develop, discontinuation of hydralazine and/or thyroxine should be considered.

Hypokalemia may develop with thiazides as with diuretics, when severe circulatory collapse, during prolonged therapy.

Hydrochlorothiazide: Serum electrolyte determination of serum electrolytes to detect hypokalemia, hypochloremia, and metabolic acidosis.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

Hydralazine: Use cautiously oral intake of diuretics, particularly in diabetic patients.

High-Risk Target Suggested In Urinary Tract Screening

Medical Tribune Report

SAN FRANCISCO—Which children are at risk for urinary tract infections? How can these infections be detected? And how should they be treated?

Dr. Patrick H. McLin, one of the speakers who discussed these questions at the annual meeting of the American Academy of Pediatrics here, described a mass screening program in which 86 of 13,148 children tested were found to be infected.

All were girls, said Dr. McLin, who estimated that 5 per cent of all girls will have trouble with urinary tract infections by the time of puberty.

The purpose of the screening, which was performed in the home by parents with a dip slide and sent back to schools for evaluation, was to determine morbidity as well as the incidence of actual or potential pyelonephritis. No pyelonephritis was detected.

Of those with infections, 44 per cent had a history of prior urinary tract disease, Dr. McLin reported. Forty per cent had symptoms of daytime wetting, frequency, urgency, or dysuria—indications that the infection was "hidden only because no one was watching."

Many Unaware of Infections

Since many of the mothers were aware of the symptoms but not of the infections, an educational effort should be made to teach mothers what is abnormal, he suggested.

Of the 86 with infections three had reflux, but no evidence of scarring was found.

Dr. McLin put the screening cost at \$21,700, or about \$1.65 per child, not counting the cost of labor, which was volunteered. The cost per infection was approximately \$250—indicating that if mass screening is to be feasible, a target population should be defined, he said.

This population should exclude boys and should include only high-risk girls in the kindergarten through sixth-grade age groups, he said. High-risk girls, he added, would include those with a high rate of absenteeism.

Dr. James E. Keeton of Jackson, Miss., said that urinary tract infections appear to be less frequent among black girls than among white and also less serious, with fewer abnormalities on intravenous pyelograms.

He also said that the incidence of reflux appears to be low and confined to preschool girls, with a high incidence of spontaneous resolution.

Dr. Joseph Y. Dwoskin of Buffalo, N.Y., observed that the infections seen by a pediatric urologist are usually more serious than those seen by a pediatrician since referrals are usually made only after two or more recurrences. The largest group of patients is in the three-to-four-year-age range,

he said, and 75 per cent are under seven years.

Unless the patients are on continuing antibiotic therapy, 50 to 65 per cent will have a recurrence within six months and 70 to 85 per cent within one year, he continued.

In one group with recurrent infections, 44 per cent had reflux and 25 per cent pyelonephritis, Dr. Dwoskin reported.

The incidence of reflux suggests that urethral manipulation should be part of the treatment for such patients, he remarked, and the incidence of pyelonephritis that investigation should be made earlier than usual. He suggested a workup after the first infection.

'At Home' Insight Into Heart Surgery Impact



Students from Stanford University Medical School visited a woman recovering from heart surgery "at home" recently for an insight into the impact that such an event has on patients and their families, emotionally and financially. Left to right: Leone McGann, Assistant Professor; Jaime Fay, student; Edith Turner, former patient; John Sanchez, student; David Kuplin, Ph.D., director of clinical social work at Stanford; and Holly Stiegman, student.

Space age microbial power BETADINE ANTISEPTICS

PHOTOGRAPH: BETHLEHEM IN ORBIT, COURTESY OF THE NATIONAL AERONAUTICS AND SPACE ADMINISTRATION.



BETADINE Skin Cleanser and **BETADINE Ointment** provide the same broad-spectrum microbial action as **BETADINE** microbicides chosen by NASA for the Skylab mission and for Apollo 11, 12/14 splashdowns. They kill gram-positive and gram-negative bacteria (including antibiotic-resistant strains), fungi, viruses, protozoa and yeasts... are virtually nonirritating and nonstinging... nonstaining to skin and natural fabrics.

BETADINE Skin Cleanser degerms the skin of patients with common pathogens, including *Staph. aureus*... helps prevent recurrence of acute inflammatory skin infections and spread of infection in acne pimples... may be used routinely for general skin hygiene.

(In the rare instance of local irritation or sensitivity, discontinue use in the individual.)

BETADINE Ointment kills pathogens in skin and wound infections... indicated in infected stasis ulcers and to help prevent infection in burns, lacerations and abrasions. Not greasy or sticky... the treated area can be bandaged.

Purdue Frederick

© 1974, The Procter & Gamble Company

DETROIT 222-2222

222-2222

DETROIT 222-2222

If there's good reason
to prescribe
for psychic tension...

Prompt action
is a good reason
to consider Valium®
(diazepam)



When, for example, despite counseling,
tension and anxiety continue to produce
distressing somatic symptoms

When your patient's somatic complaints are associated with tension and anxiety and you have tried counseling and other supportive measures alone, you may decide to prescribe psychotherapeutic medication. If you do, the question remains: which one?

Valium (diazepam) is one to consider closely. One that works promptly as an adjunct to continued supportive measures. One that generally produces significant improvement within the first few days of therapy, although some patients may require more time for a clearcut response.

Prompt action. One good reason to consider Valium.

And should you choose to prescribe Valium, you should also keep this information in mind. Valium is usually well tolerated. Patients taking Valium should be cautioned against operating dangerous machinery or driving. Therapy with Valium should normally be continued until the patient's psychic tension symptoms have been reduced to tolerable levels.

Please turn page for a summary of product information.

Valium®
(diazepam)

2-mg, 5-mg, 10-mg tablets

ROCHE

Information
to prescribers

Valium® (diazepam)

Before prescribing, please consult complete product information, a summary of which follows:

Indications: Tension and anxiety states; somatic complaints which are concomitants of emotional factors; psychoneurotic states manifested by tension, anxiety, apprehension, fatigue, depressive symptoms or agitation; symptomatic relief of acute agitation, tremor, delirium tremens and hallucinosis due to acute alcohol withdrawal; adjunctively in skeletal muscle spasm due to reflex spasm to local pathology, spasticity caused by upper motor neuron disorders, athetosis, stiff-man syndrome, convulsive disorders (not for sole therapy).

Contraindicated: Known hypersensitivity to the drug. Children under 6 months of age. Acute narrow angle glaucoma; may be used in patients with open angle glaucoma who are receiving appropriate therapy.

Warnings: Not of value in psychotic patients. Caution against hazardous occupations requiring complete mental alertness. When used adjunctively in convulsive disorders, possibility of increase in frequency and/or severity of grand mal seizures may require increased dosage of standard anti-convulsant medication; abrupt withdrawal may be associated with temporary increase in frequency and/or severity of seizures. Advise against simultaneous ingestion of alcohol and other CNS depressants. Withdrawal symptoms (similar to those with barbiturates and alcohol) have occurred following abrupt discontinuance (convulsions, tremor, abdominal and muscle cramps, vomiting and sweating). Keep addiction-prone individuals under careful surveillance because of their predisposition to habituation and dependence. In pregnancy, lactation or women of childbearing age, weigh potential benefit against possible hazard.

Precautions: If combined with other psychotropics or anticonvulsants, consider carefully pharmacology of agents employed.

drugs such as phenothiazines, narcotics, barbiturates, MAO inhibitors and other anti-depressants may potentiate its action. Usual precautions indicated in patients severely depressed, or with latent depression, or with suicidal tendencies. Observe usual precautions in impaired renal or hepatic function. Limit dosage to smallest effective amount in elderly and debilitated to preclude ataxia or oversedation.

Side Effects: Drowsiness, confusion, diplopia, hypotension, changes in libido, nausea, fatigue, depression, dysarthria, jaundice, skin rash, ataxia, constipation, headache, incontinence, changes in salivation, slurred speech, tremor, vertigo, urinary retention, blurred vision. Paradoxical reactions such as acute hyperexcited states, anxiety, hallucinations, increased muscle spasticity, insomnia, rage, sleep disturbances, stimulation have been reported; should these occur, discontinue drug. Isolated reports of neutropenia, jaundice; periodic blood counts and liver function tests advisable during long-term therapy.

Dosage: Individualize for maximum beneficial effect. **Adults:** Tension, anxiety and psychoneurotic states, 2 to 10 mg b.i.d. to q.i.d.; alcoholism, 10 mg t.i.d. or q.i.d. in first 24 hours, then 5 mg t.i.d. or q.i.d. as needed; adjunctively in skeletal muscle spasm, 2 to 10 mg t.i.d. or q.i.d.; adjunctively in convulsive disorders, 2 to 10 mg b.i.d. to q.i.d. **Geriatric or debilitated patients:** 2 to 2½ mg, 1 or 2 times daily initially, increasing as needed and tolerated. (See Precautions.) **Children:** 1 to 2½ mg t.i.d. or q.i.d. initially, increasing as needed and tolerated (not for use under 6 months).

Supplied: Valium® (diazepam) Tablets, 2 mg, 5 mg and 10 mg; bottles of 100 and 500. All strengths also available in Tel-E-Dose® packages of 100.

ROCHE
Roche Laboratories
Division of Hoffmann-La Roche Inc.
Nutley, New Jersey 07110

Wrist Prosthesis Simulates Movement of Normal Joint

Medical Tribune Report

TUSCON, ARIZ.—A completely mobile wrist prosthesis of metal and plastic, which closely resembles the bidirectional movement of the normal joint, has been designed and successfully implanted in two patients in a collaborative effort here by members of the Department of Surgery and Mechanical Engineering at the University of Arizona.

According to a preliminary report of the work by Dr. Robert G. Volz, orthopedic surgeon and Assistant Professor of Surgery at the university's College of Medicine, the replacement is not a ball-and-socket mechanism, which would cause unnatural movement, but half of a toroidal sector (shaped like a tire cut in half), fitted with an elliptical cup. Like the normal wrist, the combination allows for motion in two planes only—up-and-down (flexion/extension) and side-to-side (radial and ulnar deviation).

Both halves are held in place by methylmethacrylate. Three bones in the hand—the lunate, navicular and the head of the capitate—were resected to make room for the prosthesis.

Unlike most other joints in the body, which must withstand mostly forces of compression, the wrist must be able to take forces of distraction as well—the tendency of the joint to be pulled apart when a suitcase is lifted, for example.

The Arizona team noted that patients have been able to lift a 40-pound suitcase and squeeze a ball tightly. Although clinical data is far from complete, the musician is already back at the organ, and the collaborative team is optimistic that the replacement will permit most normal life activity.

Because of the use of methyl methacrylate cement, which is used extensively in hip-joint replacements but rarely elsewhere, the University of Arizona group has had to obtain special permission from the FDA for each operation performed so far.

Clinical data is still incomplete, the Arizona team said, but the operation may be indicated for many patients with crushed or deformed wrists, and for persons with rheumatoid arthritis of the wrist without metacarpal, phalangeal, or interphalangeal involvement. The prosthesis would probably not be useful in replacing the wrist of an arthritic patient with appreciable hand or finger involvement.

One of the first patients to receive

the operation was a printer and part-time organist whose left hand was badly crushed in an accident. The two sections of the prosthesis were cemented to the radius on one end and the bones of the second and third fingers on the other. The metal portion is made of Vitallium, an alloy of cobalt chromium, which is not rejected by the body.

Both halves are held in place by methylmethacrylate. Three bones in the hand—the lunate, navicular and the head of the capitate—were resected to make room for the prosthesis.

Unlike most other joints in the body, which must withstand mostly forces of compression, the wrist must be able to take forces of distraction as well—the tendency of the joint to be pulled apart when a suitcase is lifted, for example.

The Arizona team noted that patients have been able to lift a 40-pound suitcase and squeeze a ball tightly. Although clinical data is far from complete, the musician is already back at the organ, and the collaborative team is optimistic that the replacement will permit most normal life activity.

Because of the use of methyl methacrylate cement, which is used extensively in hip-joint replacements but rarely elsewhere, the University of Arizona group has had to obtain special permission from the FDA for each operation performed so far.

Assisting in the design of the prosthesis were Drs. Marvin D. Martin, Professor of Mechanical Engineering and Michael J. Pitt of the department of radiology. Mr. Richard Perry, a student in the Medical College, was also part of the team.

Disappointment was voiced that a proposed abortion law, now pending in the Knesset, Israel's Parliament, provided that committees would be set up to consider requests for abortion.

Several speakers pointed out that this favored the rich and well-to-do.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Despite brave talk of keeping Federal disbursements down to \$300 billion a year, they are headed closer to \$400 billion.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for

Tel Aviv—"Abortion on demand" was urged here recently by the National Society of Public Health and the Society of Public Health Physicians at a joint meeting.

Government actions calling for



What a difference a day can make

Your counsel and reassurance—Ritalin.

A logical first step in treating mild depression,* and often all that's needed to bring quick symptomatic relief.

Indeed, your patient may be

grin to feel better within hours—her spirits boosted, her mood brightened. A single prescription may be all that's needed.

Ritalin is usually well tolerated even by older or convalescent patients. Note, however, that it is not indicated in the more severe depressions.

But whenever depression is mild, think of Ritalin—so your patient has a better chance of waking up to a brighter tomorrow.

Ritalin[®]
(methylphenidate)
acts quickly to relieve symptoms
in mild depression

*This drug has been evaluated as possibly effective for this indication. See brief prescribing information.

Ritalin[®] hydrochloride
(methylphenidate hydrochloride)
TABLETS

INDICATION
Based on a review of this drug by the National Academy of Sciences-National Research Council and/or other information, FDA has classified the indication as "possibly" effective. Mild depression. Final classification of the indication will depend on further investigation.

CONTRAINdications
Marked anxiety, tension, and agitation, since Ritalin may aggravate these symptoms. Also contraindicated in patients known to be hypersensitive to the drug, and in patients with phimosis.

WARNING
Ritalin should not be used in children under six years, since safety and efficacy in this age group have not been established.

Sufficient data on safety and efficacy of long-term use of Ritalin in children with minimal brain dysfunction are not yet available. Although a causal relationship has not been established, an association of growth (i.e., weight gain and/or height) has been reported to the long-term use of stimulants in children. Therefore, children requiring long-term therapy should be carefully monitored.

Ritalin should not be used for severe depression of either exogenous or endogenous origin or for the prevention of normal fatigue states.

Ritalin may lower the convulsive threshold in patients with or without seizures, even without prior EEG abnormalities, even in absence of seizures. Safe concomitant use of anticonvulsants and Ritalin has not been established. If seizures occur, Ritalin should be discontinued.

Use cautiously in patients with hypertension. Blood pressure should be monitored at appropriate intervals in patients taking Ritalin, especially those with hypertension.

Drug Interactions

Ritalin may decrease the hypotensive effect of guanethidine. Use cautiously with pressor agents and MAO inhibitors. Ritalin may inhibit the metabolism of amine and captopril, and may potentiate propranolol, diphenhydramine, primaquine, chloroquine, and cyclic antidepressants. Amphetamine, decongestants, downward the age of 12 treatment of these drugs may be required when given concomitantly with Ritalin.

Usage in Pregnancy

Adverse reactions in both mother to establish safety of Ritalin during pregnancy have not been conducted. Therefore, until more information is available, Ritalin should not be prescribed for women of childbearing age unless, in the opinion of the physician, the potential benefits outweigh the possible risks.

Drug Dependence

Ritalin should be given cautiously to come chronically medicated patients, such as those with a history of drug dependence or alcoholism, because in such patients, may increase damage on their own initiative.

Chronically obese patients tend to marked tolerance and begin dependence with continued use. To alleviate this risk, frank psychiatric notches, eating disorder, especially with parenteral abuse. Cautious supervision is required during drug withdrawal, since severe depression as well as the effects of chronic overuse may be increased.

"The hazard to young athletes, such as high-school football players who begin training in the summer, is that a heavy daily exercise schedule can quickly produce cumulative potassium losses, mostly from sweating," he said.

Although the average daily dietary intake of potassium is about 75-100 mEq, hard exercise on a hot day can lead to a loss of 100 mEq, through sweating alone, Dr. Knochel explained, and if the loss through urine of about 50-60 mEq, is added an over-all loss quickly accumulates.

"We did one study in which six army recruits, training in summer heat, were found to have a serious potassium deficit by the 11th day of training," he related. "By contrast, 16 other subjects studied in cool weather in identical fashion—using radioactively tagged potassium—did not become deficient."

Effect on Blood Supply

Ordinarily, Dr. Knochel said, potassium released by muscle cells during contraction acts on local blood vessels to increase blood supply to the exercised muscle.

In experimental studies, he detected a significant increase of blood potassium levels and blood flow when specific muscles of normal dogs were exercised.

Average dose is 20 to 30 mg daily. Some patients may require 40 to 60 mg daily. In others, 10 to 15 mg daily will be adequate. The few patients who are unable to sleep if medication is taken late in the day should take the last dose, before 9 p.m.

HOW SUPPLIED

Tablets, 20 mg (peach, scored); bottles of 100, and 1000.

Tablets, 10 mg (peach green, scored); bottles of 100, 500, 1000 and Accu-pak blister units of 100.

Tablets, 5 mg (peach yellow); bottles of 100, 500 and 1000.

Consult complete product literature before prescribing.

CIBA Pharmaceutical Company

Division of CIBA-GEIGY Corporation

Summit, New Jersey 07901

© 1975 CIBA-GEIGY Corporation

50 mg (peach, scored); bottles of 100, and 1000.

Tablets, 10 mg (peach green, scored); bottles of 100, 500, 1000 and Accu-pak blister units of 100.

Tablets, 5 mg (peach yellow); bottles of 100, 500 and 1000.

Consult complete product literature before prescribing.

CIBA Pharmaceutical Company

Division of CIBA-GEIGY Corporation

Summit, New Jersey 07901

© 1975 CIBA-GEIGY Corporation

50 mg (peach, scored); bottles of 100, and 1000.

Tablets, 10 mg (peach green, scored); bottles of 100, 500, 1000 and Accu-pak blister units of 100.

Tablets, 5 mg (peach yellow); bottles of 100, 500 and 1000.

Consult complete product literature before prescribing.

CIBA Pharmaceutical Company

Division of CIBA-GEIGY Corporation

Summit, New Jersey 07901

© 1975 CIBA-GEIGY Corporation

50 mg (peach, scored); bottles of 100, and 1000.

Tablets, 10 mg (peach green, scored); bottles of 100, 500, 1000 and Accu-pak blister units of 100.

Tablets, 5 mg (peach yellow); bottles of 100, 500 and 1000.

Consult complete product literature before prescribing.

CIBA Pharmaceutical Company

Division of CIBA-GEIGY Corporation

Summit, New Jersey 07901

© 1975 CIBA-GEIGY Corporation

50 mg (peach, scored); bottles of 100, and 1000.

Tablets, 10 mg (peach green, scored); bottles of 100, 500, 1000 and Accu-pak blister units of 100.

Tablets, 5 mg (peach yellow); bottles of 100, 500 and 1000.

Consult complete product literature before prescribing.

CIBA Pharmaceutical Company

Division of CIBA-GEIGY Corporation

Summit, New Jersey 07901

© 1975 CIBA-GEIGY Corporation

50 mg (peach, scored); bottles of 100, and 1000.

Tablets, 10 mg (peach green, scored); bottles of 100, 500, 1000 and Accu-pak blister units of 100.

Tablets, 5 mg (peach yellow); bottles of 100, 500 and 1000.

Consult complete product literature before prescribing.

CIBA Pharmaceutical Company

Division of CIBA-GEIGY Corporation

Summit, New Jersey 07901

© 1975 CIBA-GEIGY Corporation

50 mg (peach, scored); bottles of 100, and 1000.

Tablets, 10 mg (peach green, scored); bottles of 100, 500, 1000 and Accu-pak blister units of 100.

Tablets, 5 mg (peach yellow); bottles of 100, 500 and 1000.

Consult complete product literature before prescribing.

CIBA Pharmaceutical Company

Division of CIBA-GEIGY Corporation

Summit, New Jersey 07901

© 1975 CIBA-GEIGY Corporation

50 mg (peach, scored); bottles of 100, and 1000.

Tablets, 10 mg (peach green, scored); bottles of 100, 500, 1000 and Accu-pak blister units of 100.

Tablets, 5 mg (peach yellow); bottles of 100, 500 and 1000.

Consult complete product literature before prescribing.

CIBA Pharmaceutical Company

Division of CIBA-GEIGY Corporation

Summit, New Jersey 07901

© 1975 CIBA-GEIGY Corporation

50 mg (peach, scored); bottles of 100, and 1000.

Tablets, 10 mg (peach green, scored); bottles of 100, 500, 1000 and Accu-pak blister units of 100.

Tablets, 5 mg (peach yellow); bottles of 100, 500 and 1000.

Consult complete product literature before prescribing.

CIBA Pharmaceutical Company

Division of CIBA-GEIGY Corporation

Summit, New Jersey 07901

© 1975 CIBA-GEIGY Corporation

50 mg (peach, scored); bottles of 100, and 1000.

Tablets, 10 mg (peach green, scored); bottles of 100, 500, 1000 and Accu-pak blister units of 100.

Tablets, 5 mg (peach yellow); bottles of 100, 500 and 1000.

Consult complete product literature before prescribing.

CIBA Pharmaceutical Company

Division of CIBA-GEIGY Corporation

Summit, New Jersey 07901

© 1975 CIBA-GEIGY Corporation

50 mg (peach, scored); bottles of 100, and 1000.

Tablets, 10 mg (peach green, scored); bottles of 100, 500, 1000 and Accu-pak blister units of 100.

Tablets, 5 mg (peach yellow); bottles of 100, 500 and 1000.

Consult complete product literature before prescribing.

CIBA Pharmaceutical Company

Division of CIBA-GEIGY Corporation

Summit, New Jersey 07901

© 1975 CIBA-GEIGY Corporation

50 mg (peach, scored); bottles of 100, and 1000.

Tablets, 10 mg (peach green, scored); bottles of 100, 500, 1000 and Accu-pak blister units of 100.

Tablets, 5 mg (peach yellow); bottles of 100, 500 and 1000.

Consult complete product literature before prescribing.

CIBA Pharmaceutical Company

Division of CIBA-GEIGY Corporation

Summit, New Jersey 07901

© 1975 CIBA-GEIGY Corporation

50 mg (peach, scored); bottles of 100, and 1000.

Tablets, 10 mg (peach green, scored); bottles of 100, 500, 1000 and Accu-pak blister units of 100.

Tablets, 5 mg (peach yellow); bottles of 100, 500 and 1000.

Consult complete product literature before prescribing.

CIBA Pharmaceutical Company

Division of CIBA-GEIGY Corporation